mRNA was detected in samples from liver, kidney, pancreas, stomach, thyroid, prostate and with multiple human tissues blots (CLONTECH) using a full-length human hepsin cDNA. Hepsin Fig. 1. Northern analysis of hepsin mRNA in human tissues. Northern hybridization was performed



heart brain placenta lung liver skeletal muscle kidney pancreas stomach thyroid spinal cord lymph node trachea adrenal gland bone marrow spleen thymus prostate testis ovary small intestine colon peripheral blood leukocyte

# Relative Level of Hepsin mRNA Expression



Fig. 2. Taqman analysis of hepsin mRNA expression in human tissues. Taqman-based quantitative PCR analysis of hepsin mRNA expression was performed using two sets of hepsin-specific primers and kidney. (Primer-Probes 1 and 2). High levels of hepsin mRNA expression were detected in samples from liver

and prostate cancer tissues. Northern blot was prepared and hybridized with a human hepsin cDNA ratio of hepsin/GAPDH mRNA was  $\sim$ 6-fold higher in the advanced cancer than in normal prostate, probe. Hepsin mRNA expressed was significantly higher in an advanced prostate cancer sample. The Fig. 3. Hepsin mRNA expression in prostate cancer. mRNA was isolated from normal prostate, BPH BPH or primary prostate cancer. Hepsin/GAPDH (fold increase) ယ Ŋ Prostate Normal Prostate Grade 3 Grade 4 hepsin

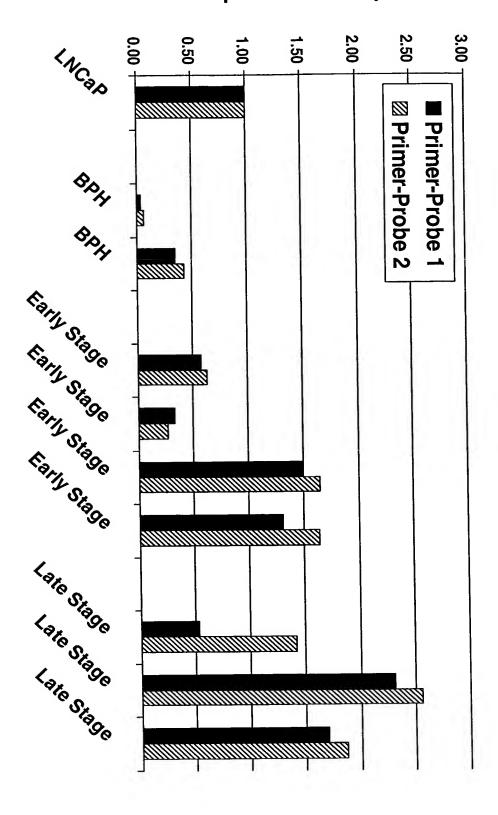
Normal

Normal

**BPH** 

Primary cancer

Advanced cancer

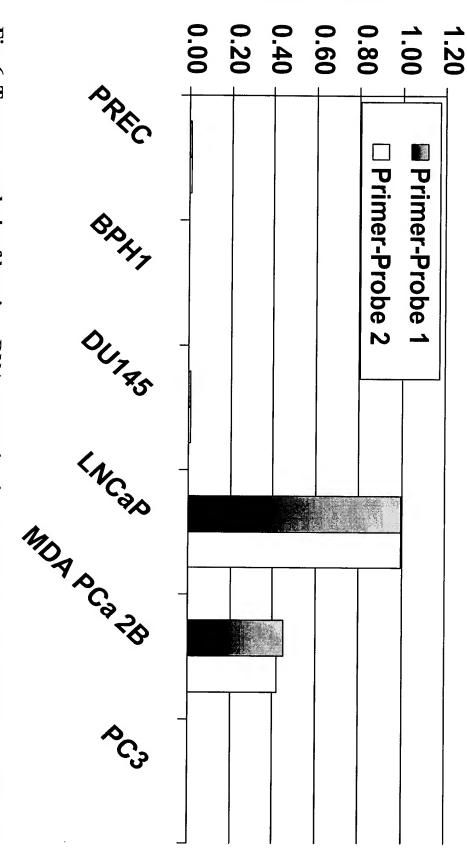


performed using two set of primers specific for human hepsin (Primer-Probes 1 and 2). Modest to more dramatic Figure 4. Taqman analysis of hepsin mRNA expression in prostate tissue. mRNA samples were prepared from increases of hepsin mRNA expression were detected in prostate cancer samples. LNCaP cells, BPH and prostate cancer (Gleason grade 3 and 4) tissues. Quantitative PCR analysis was

Fig. 5. Northern analysis of hepsin mRNA expression in prostate-derived cell lines. mRNA was with ethdium bromide to show equal sample loading. isolated from prostate-derived cell lines, LNCaP, PC3, MDA PCa2b, BPH-1, DU145, PZ HPV7, in LNCaP, MDA Pca 2b and HepG2 cells (upper panel). As a control, the agarose gel was stained CA HPV10 and a control cell line, HepG2. Northern analysis detected hepsin mRNA expression

W HepG2 LNCaP PC3 MDA PCa 2b **BPH-1 DU145** PZ HPV7 CA HPV10

### Relative Expression of Hepsin mRNA



LNCaP and MDA Pca 2B cells. hepsin specific primers (Primer-Probes 1 and 2). Hepsin mRNA expression was detected in isolated from prostate-derived cell lines. Taqman analysis was performed using two sets of Fig. 6. Taqman analysis of hepsin mRNA expression in prostate-derived cell lines. mRNA was

Hepsin/GAPDH

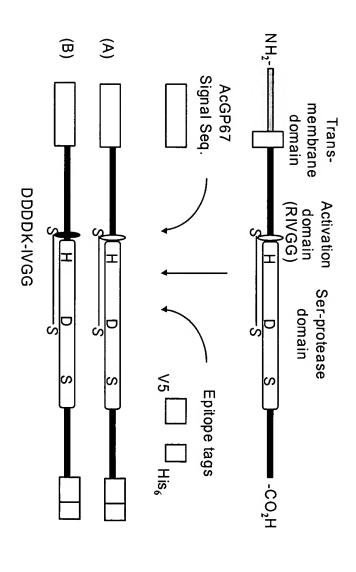
**B**.

(fold increase) ယ္မ 55 vehicle veh**is** 22 hr 14.54u

Hepsin

samples were isolated. Northern blots were hybridized with a human hepsin cDNA probe and re-probed with a GAPDH cDNA. Hepsin mRNA expression was significantly enhanced in Fig. 7. Up-regulation of hepsin mRNA expression in LNCaP cells by dihydrotestosterone (DHT). LNCaP cells treated with DHT for 24 and 72 hours. LNCaP cells were treated with DHT (10 nM) and harvested at indicted time points. Total RNA

DHT



sequence for Arg 162. The mutation will allow hepsin to be activated by enterokinase. sequence coding for an enterokinase recognition peptide, DDDDK, is inserted to replace the coding 5' end and a sequence coding for a V5 epitope and a His6 tag at the 3' end. In construct B, a of hepsin is cloned into vector pAcGP67 that contains a sequence coding for a signal peptide at the hepsin protein is shown on top. To express soluble wild-type hepsin (A), the extracellular domain Fig. 8. Expression of soluble wild-type and mutant hepsins. A schematic presentation of human

# HG9 (1-7) pIRESpuro2W/hepEK\_k

1	GACGGATCGG	GAGATCTCCC	GATCCCCTAT	GGTCGACTCT	CAGTACAATC
					GTCATGTTAG
51	TGCTCTGATG	CCGCATAGTT	AAGCCAGTAT	CTGCTCCCTG	CTTGTGTGTT
	ACGAGACTAC	GGCGTATCAA	TTCGGTCATA	GACGAGGGAC	GAACACACAA
101	GGAGGTCGCT	GAGTAGTGCG	CGAGCAAAAT	TTAAGCTACA	ACAAGGCAAG
	CCTCCAGCGA	CTCATCACGC	GCTCGTTTTA	AATTCGATGT	TGTTCCGTTC
151					GCGTTTTGCG
	CGAACTGGCT	GTTAACGTAC	TTCTTAGACG	AATCCCAATC	CGCAAAACGC
201			CAGATATACG		
	GACGAAGCGC	TACATGCCCG	GTCTATATGC	GCAACTGTAA	СТААТААСТС
251					AGCCCATATA
	ATCAATAATT	ATCATTAGTT	AATGCCCCAG	TAATCAAGTA	TCGGCTATAT
301			CTTACGGTAA		
	ACCTCAAGGC	GCAATGTATT	GAATGCCATT	TACCGGGGGG	ACCCACTCCC
351			GACGTCAATA		
	GGGTTGCTGG	GGGCGGGTAA	CTGCAGTTAT	TACTGCATAC	AACCCTATAGT
401			ATTGACGTCA		
	TTGCGGTTAT	CCCTGAAAGG	TAACTGCAGT	TACCCACCTC	ATTACGGT
451			CATCAAGTGT		
	TTTGACGGGT	GAACCGTCAT	GTAGTTCACA	TACTATACCC	TTCATCCCCC
501			TAAATGGCCC		
	GGATAACTGC	AGTTACTGCC	ATTTACCGGG	CGGACCGTAA	TACCCCTCAT
551			CTACTTGGCA		
	GTACTGGAAT	ACCCTGAAAG	GATGAACCGT	CATCTACATC	CATATTAGTCA
601			CGGTTTTGGC		
	AGCGATAATG	GTACCACTAC	GCCAAAACCG	TCATCTACTT	ACCCCCACCT
651			ATTTCCAAGT		
	ATCGCCAAAC	TGAGTGCCCC	TAAAGGTTCA	GAGGTGGGGT	AACTCCACTT
701			AAAATCAACG		
	ACCCTCAAAC	AAAACCGTGG	TTTTAGTTGC	CCTGAAAGGT	TTTACAGCAT
751			CAAATGGGCG		
	TGTTGAGGCG	GGGTAACTGC	GTTTACCCGC	CATCCGCACA	TGCCACCCTC
801			CTGGCTAACT		
	CAGATATATT	CGTCTCGAGA	GACCGATTGA	TCTCTTGGGT	GACGAATGAC
851			CTCACTATAG		
	CGAATAGCTT	TAATTATGCT	GAGTGATATC	CCTCTGGGTT	CGAACCATGG
+3				TDTL	
901	GAGCTCGGAT	CGATATCGCC	ACCATGGAGA	CAGACACACT	CCTGCTATGG
	CTCGAGCCTA	${\tt GCTATAGCGG}$	TGGTACCTCT	GTCTGTGTGA	GGACGATACC
+3		L W V P			P D R S
951	GTACTGCTGC	TCTGGGTTCC	AGGTTCCACT	GGTGACGCTC	CGGACAGGAG
	CATGACGACG	AGACCCAAGG	TCCAAGGTGA	CCACTGCGAG	GCCTGTCCTC
+3	SDQE	PLY	P V Q V	S S A	DARL
1001	TGACCAGGAG	CCGCTGTACC	CAGTGCAGGT	CAGCTCTGCG	GACGCTCGGC
	ACTGGTCCTC	GGCGACATGG	GTCACGTCCA	GTCGAGACGC	CTGCGAGCCG

+3 · L	M V F D K T E G T W R L L C S S
	TO THE STREET, THE ACADELIC GAAGGGACGT GGCGGCTGCT GTGCTCCTCG
1051 TO	CTACCAGAA ACTGTTCTGC CTTCCCTGCA CCGCCGACGA CACGAGGAGG
.2 1	R S N A R V A G L S C E E M G I L
1101 C	GCTCTAACG CCAGGGTAGC CGGACTCAGC TGCGAGGAGA TGGGCTTCCT
G	CGAGATTGC GGTCCCATCG GCCTGAGTCG ACGCTCCTCT ACCCGAAGGT
+3 ·L	RALTHS ELD V R 1 A CONTROL
1151 C	AGGGCACTG ACCCACTCCG AGCTGGACGT GCGAACGGCG GGCGCCAATG
	CAGGGCACTG ACCCACTOOS TOO TOO TOO TOO TOO TOO TOO TOO TOO
+3	G T S G F F C V D E G R E  GCACGTCGGG CTTCTTCTGT GTGGACGAGG GGAGGCTGCC CCACACCCAG  GCACGTCGGG CTTCTTCTGT GTGGACGAGG GGAGGCTGCC CCTCTCGGTC
1201	GCACGTCGGG CTTCTTCTGT GTGGACGACG GCTCCGACGG GGTGTGGGTC
	R L L E V I S V C D C P R G R F L.
+3	COORDINATE ACCURATOR CONTINUE TO THE ACCURAGE CONTINUE TO THE ACCURACY
1251	TCCGACGACC TCCAGTAGAG GCACACACTA ACGGGGTCTC CGGCAAAATT
	LAALCQDCGRRKLPVDD
1301	GGCCGCCATC TGCCAAGACT GTGGCCGCAG GAAGCTGCCC GTGGACGACG
	CCGGCGGTAG ACGGTTCTGA CACCGGCGTC CTTCGACGGG CACCTGCTGC
+3	DDKIVGGRDI3
1351	ACGACAAGAT CGTGGGAGGC CGGGACACCA GCTTGGGCCG GTGGCCGTGG TGCTGTTCTA GCACCCTCCG GCCCTGTGGT CGAACCCGGC CACCGGCACC
	a . p v p g A H L C G G S L L
+3	THE STOREGE THE COUNTY TO THE TOTAL TOTAL TOTAL COUNTY TO THE TOTAL COUNTY TO THE TOTAL TO
1401	CTTCACTCGC AAGCGATACT ACCTCGTGTG GAGACACCCC CTAGGGACACCT
+3	ISG D W V L T A A H C F P E R R R
1451	CTCCGGGGAC TGGGTGCTGA CAGCCGCCCA CTGCTTCCCG GAGCGGAACC
	GAGGCCCCTG ACCCACGACT GTCGGCGGGT GACGAAGGGC CTCGCCTTGG
+3	RVISRWRVFAGA
1501	GGGTCCTGTC CCGATGGCGA GTGTTTGCCG GTGCCGTGGC CCAGGCCTCT CCCAGGACAG GGCTACCGCT CACAAACGGC CACGGCACCG GGTCCGGAGA
+3	P H G L Q L G V Q A  CCCCACGGTC TGCAGCTGGG GGTGCAGGCT GTGGTCTACC ACGGGGGCTA  CCCCACGGTC TGCAGCTGGG GGTGCAGGCT GTGGCCCCGAT
1551	CCCCTCCCAG ACGTCGACCC CCACGTCCGA CACCAGAIGG IGCCCCCAI
+3	VIPERDPNSEENSNUTT
1601	TCTTCCCTTT CGGGACCCCA ACAGCGAGGA GAACAGCAAC GATATTGCCC
1001	AGAAGGAAA GCCCTGGGGT TGTCGCTCCT CTTGTCGTTG CTATAACGG
+3	LVHLSSPLPLTEYIQPV
1651	TGGTCCACCT CTCCAGTCCC CTGCCCCTCA CAGAATACAT CCAGCCTGTG
	ACCAGGTGGA GAGGTCAGGG GACGGGGAGT GTCTTATGTA GGTCGGACAC
+3	C L P A A G Q A L V D G K  TGCCTCCCAG CTGCCGGCCA GGCCCTGGTG GATGGCAAGA TCTGTACCGT  TGCCTCCCAG CTGCCGGCCA GGCCCTGGTG GATGGCAAGA TCTGTACCGCA
1701	TGCCTCCCAG CTGCCGGCCA GGCCCTGGTG GATGGCATTCT AGACATGGCA ACGGAGGGTC GACGGCCGGT CCGGGACCAC CTACCGTTCT AGACATGGCA
+3	VIGWGNIQYYGQQAAG
	THE THE GOOD ACACOC ACTACTATES CCAACAGGCC GGGGTACTCC
1751	GACGGGCTGG GGCAACACGC NOTIFICATACC GGTTGTCCGG CCCCATGAGG

+3	QEARVPIISND V CN G A D
1801	AGGAGGCTCG AGTCCCCATA ATCAGCAATG ATGTCTGCAA TGGCGCTGAT
1001	TCCTCCGAGC TCAGGGGTAT TAGTCGTTAC TACAGACGTT ACCGCGACTA
+3	FYGNQIKPKM FCAGYPE
1851	TTCTATGGAA ACCAGATCAA GCCCAAGATG TTCTGTGCTG GCTACCCCGA
	AAGATACCTT TGGTCTAGTT CGGGTTCTAC AAGACACGAC CGATGGGGCT
+3	EGGIDACQGDSGGPFVCE
1901	GGGTGGCATT GATGCCTGCC AGGGCGACAG CGGTGGTCCC TTTGTGTGTG
	CCCACCGTAA CTACGGACGG TCCCGCTGTC GCCACCAGGG AAACACACAC
+3	EDS I SRT PRW R L C G I V S
1951	AGGACAGCAT CTCTCGGACG CCACGTTGGC GGCTGTGTGG CATTGTGAGT
_	TCCTGTCGTA GAGAGCCTGC GGTGCAACCG CCGACACACC GTAACACTCA W G T G C A L A Q K P G V Y T K V
+3	Wdiad
2001	TGGGGCACTG GCTGTGCCCT GGCCCAGAAG CCAGGCGTCT ACACCAAAGT ACCCCGTGAC CGACACGGGA CCGGGTCTTC GGTCCGCAGA TGTGGTTTCA
+3	·V S D F R E W I F Q A I K T H S E A
	CAGTGACTTC CGGGAGTGGA TCTTCCAGGC CATAAAGACT CACTCCGAAG
2051	GTCACTGAAG GCCCTCACCT AGAAGGTCCG GTATTTCTGA GTGAGGCTTC
+3	ASGMVTQLEFGKPIPNP
2101	CCAGCGGCAT GGTGACCCAG CTCGAATTCG GTAAGCCTAT CCCTAACCCT
2101	GGTCGCCGTA CCACTGGGTC GAGCTTAAGC CATTCGGATA GGGATTGGGA
+3	LLGLDSTRTGHHHHH
2151	CTCCTCGGTC TCGATTCTAC GCGTACCGGT CATCATCACC ATCACCATTG
	GAGGAGCCAG AGCTAAGATG CGCATGGCCA GTAGTAGTGG TAGTGGTAAC
2201	AGTTTAAAGC GGCCGCATAG ATAACTGATC CAGTGTGCTG GAATTAATTC
	TCAAATTTCG CCGGCGTATC TATTGACTAG GTCACACGAC CTTAATTAAG
2251	GCTGTCTGCG AGGGCCAGCT GTTGGGGTGA GTACTCCCTC TCAAAAGCGG
	CGACAGACGC TCCCGGTCGA CAACCCCACT CATGAGGGAG AGTTTTCGCC
2301	GCATGACTTC TGCGCTAAGA TTGTCAGTTT CCAAAAACGA GGAGGATTTG
	CGTACTGAAG ACGCGATTCT AACAGTCAAA GGTTTTTGCT CCTCCTAAAC
2351	ATATTCACCT GGCCCGCGGT GATGCCTTTG AGGGTGGCCG CGTCCATCTG
	TATAAGTGGA CCGGGCGCCA CTACGGAAAC TCCCACCGGC GCAGGTAGAC
2401	GTCAGAAAAG ACAATCTTTT TGTTGTCAAG CTTGAGGTGT GGCAGGCTTG CAGTCTTTTC TGTTAGAAAA ACAACAGTTC GAACTCCACA CCGTCCGAAC
0.454	
2451	AGATCTGGCC ATACACȚTGA GTGACAATGA CATCCACTTT GCCTTTCTCT TCTAGACCGG TATGTGAACT CACTGTTACT GTAGGTGAAA CGGAAAGAGA
0501	The state of the s
2501	GGTGTCCACA GGTGAGGGTC CAGGTTGACG TCCAGCTCGT ACGTAGATCC
2551	· · · · · · · · · · · · · · · · · · ·
2331	CGCCGGTTAA GGCGGGAGA GGGAGGGGGG GGGGATTGCA ATGACCGGCT
2601	
2001	TCGGCGAACC TTATTCCGGC CACACGCAAA CAGATATACA CTAAAAGGTC
2651	CONTRACTOR OF THE CONTRACTOR O
2001	GTATAACGGC AGAAAACCGT TACACTCCCG GGCCTTTGGA CCGGGACAGA
2701	TCTTGACGAG CATTCCTAGG GGTCTTTCCC CTCTCGCCAA AGGAATGCA
	AGAACTGCTC GTAAGGATCC CCAGAAAGGG GAGAGCGGTT TCCTTACGT

FIGURE 9-3

2751		ATGTCGTGAA TACAGCACTT			
2801		TCTGTAGCGA			
2001		AGACATCGCT			
2851	GCGACAGGTG	CCTCTGCGGC	CAAAAGCCAC	GTGTATAAGA	TACACCTGCA
	CGCTGTCCAC	GGAGACGCCG	GTTTTCGGTG	CACATATTCT	ATGTGGACGT
2901	AAGGCGGCAC	AACCCCAGTG	CCACGTTGTG	AGTTGGATAG	TTGTGGAAAG
		TTGGGGTCAC			
2951		CTCTCCTCAA			
2,7,5,1		GAGAGGAGTT			
3001		CCATTGTATG			
3001		GGTAACATAC			
2054					
3051		TTTAGTCGAG AAATCAGCTC			
3101		GTTTTCCTTT			
		CAAAAGGAAA			
3151		CGACCTTCCA			
		GCTGGAAGGT			
3201		CGACGTCCCC			
	GGTGGGCGCT	GCTGCAGGGG	GCCCGGCATG	CGTGGGAGCG	GCGGCGCAAG
3251					GCCACATCGA
	CGGCTGATGG	GGCGGTGCGC	GGTGTGGCAG	CTGGGCCTGG	CGGTGTAGCT
3301	GCGGGTCACC	GAGCTGCAAG	AACTCTTCCT	CACGCGCGTC	GGGCTCGACA
	CGCCCAGTGG	CTCGACGTTC	TTGAGAAGGA	GTGCGCGCAG	CCCGAGCTGT
3351	TCGGCAAGGT	GTGGGTCGCG	GACGACGGCG	CCGCGGTGGC	GGTCTGGACC
	AGCCGTTCCA	CACCCAGCGC	CTGCTGCCGC	GGCGCCACCG	CCAGACCTGG
3401	ACGCCGGAGA	GCGTCGAAGC	GGGGGCGGTG	TTCGCCGAGA	TCGGCCCGCG
					AGCCGGGCGC
3451					CAGATGGAAG
7471					GTCTACCTTC
3501					CCTGGCCACC
3301					GGACCGGTGG
2551					GCGCCGTCGT
3551					CGCGGCAGCA
3601					GCCTTCCTGG GCGAAGGACC
3651					CGGCTTCACC
					GCCGAAGTGG
3701					GGTGCATGAC
					CCACGTACTG
3751					CGCCCGACCG
					GCGGGCTGGC
3801					CGGGCGGCCC
	TTTCCTCGCC	G TGCTGGGGTA	CCGAGGCTGG	CTTCGGCTGC	GCCCGCCGGG
3851					ACTCGAGATC
	GCGGCTGGG	G CGTGGGCGGG	GGCTCCGGGT	GGCTGAGAT	TGAGCTCTAG

3901		CTCTGGATTA GAGACCTAAT		
3951		GCTCCTTTTA CGAGGAAAAT		
4001		TATTGCTTCC ATAACGAAGG		
4051		TGCTGTCTCT ACGACAGAGA		
4101		GTGTGCACTG CACACGTGAC		
4151		CACCTGTCAG GTGGACAGTC		
4201		CGGCGGAACT GCCGCCTTGA		
4251		CTGTTGGGCA GACAACCCGT		
4301		TCCTTGGCTG AGGAACCGAC		
4351		TCTGCTACGT AGACGATGCA		
4401		CTGCTGCCGG GACGACGGCC		
4451		GAGTCGGATC CTCAGCCTAG		
4501		GCTGATCAGC CGACTAGTCG		
4551	- <del>-</del>	CCCTCCCCG GGGAGGGGGC		
4601		TTCCTAATAA AAGGATTATT		TTGTCTGAGT AACAGACTCA
4651		CTATTCTGGG GATAAGACCC		GCAAGGGGGA CGTTCCCCCT
4701		GACAATAGCA CTGTTATCGT		
4751		GGAAAGAACC CCTTTCTTGG		CTAGTTGTGG GATCAACACC
4801				CCGTCGACCT GGCAGCTGGA
4851				TGTGTGAAAT ACACACTTTA
4901				GCATAAAGTG CGTATTTCAC
4951				ATTGCGTTGC TAACGCAACG
5001				GCTGCATTAA CCGACGTAATT

FIGURE 9-5

5051	TO A ATOCOCO	AACGCGCGGG	CACACCCCC	MMCCCM A MMC	acacamamma
3031		TTGCGCGCCC			
5101		CTCACTGACT			
3101		GAGTGACTGA			
5151		TCACTCAAAG			
		AGTGAGTTTC			
5201	GATAACGCAG	GAAAGAACAT	GTGAGCAAAA	GGCCAGCAAA	AGGCCAGGAA
	CTATTGCGTC	CTTTCTTGTA	CACTCGTTTT	CCGGTCGTTT	TCCGGTCCTT
5251	CCGTAAAAAG	GCCGCGTTGC	TGGCGTTTTT	CCATAGGCTC	CGCCCCCTG
	${\tt GGCATTTTTC}$	CGGCGCAACG	ACCGCAAAAA	${\tt GGTATCCGAG}$	GCGGGGGGAC
5301	ACGAGCATCA	CAAAAATCGA	CGCTCAAGTC	AGAGGTGGCG	AAACCCGACA
	TGCTCGTAGT	GTTTTTAGCT	GCGAGTTCAG	TCTCCACCGC	TTTGGGCTGT
5351		GATACCAGGC			
		CTATGGTCCG			
5401		ACCCTGCCGC			
5 4 F 4		TGGGACGGCG			
5451		GGCGCTTTCT			
5501		CCGCGAAAGA			
2201		TTCGCTCCAA AAGCGAGGTT			
5551		TGCGCCTTAT			
3331		ACGCGGAATA		· · · · · · · · · · · · · · · · · · ·	
5601		CTTATCGCCA			
		GAATAGCGGT			<del>-</del>
5651	AGAGCGAGGT	ATGTAGGCGG	TGCTACAGAG	TTCTTGAAGT	GGTGGCCTAA
	TCTCGCTCCA	TACATCCGCC	ACGATGTCTC	AAGAACTTCA	CCACCGGATT
5701	CTACGGCTAC	ACTAGAAGGA	CAGTATTTGG	TATCTGCGCT	CTGCTGAAGC
	GATGCCGATG	TGATCTTCCT	GTCATAAACC	ATAGACGCGA	GACGACTTCG
5751	CAGTTACCTT	CGGAAAAAGA	GTTGGTAGCT	CTTGATCCGG	CAAACAAACC
		GCCTTTTTCT			
5801		GCGGTGGTTT			
		CGCCACCAAA			
5851		TCTCAAGAAG AGAGTTCTTC			
5901					
3901					GAGATTATCA CTCTAATAGT
5951		TCACCTAGAT			
3331					CAAAATTTAG
6001					CAATGCTTAA
					GTTACGAATT
6051					ATCCATAGTT
	AGTCACTCCG	TGGATAGAGT	CGCTAGACAG	ATAAAGCAAG	TAGGTATCAA
6101	GCCTGACTCC	CCGTCGTGTA	GATAACTACG	ATACGGGAGG	GCTTACCATC
				,	CGAATGGTAG
6151					CCGGCTCCAG
	ACCGGGGTCA	CGACGTTACT	ATGGCGCTCT	GGGTGCGAGT	GGCCGAGGTC

## FIGURE 9-6

6201	ATTTATCAGC	AATAAACCAG	CCAGCCGGAA	GGGCCGAGCG	CAGAAGTGGT
	TAAATAGTCG	TTATTTGGTC	GGTCGGCCTT	CCCGGCTCGC	GTCTTCACCA
6251	CCTGCAACTT	TATCCGCCTC	CATCCAGTCT	ATTAATTGTT	GCCGGGAAGC
	GGACGTTGAA	ATAGGCGGAG	GTAGGTCAGA	TAATTAACAA	CGGCCCTTCG
6301	TAGAGTAAGT	AGTTCGCCAG	${\tt TTAATAGTTT}$	$\tt GCGCAACGTT$	GTTGCCATTG
	ATCTCATTCA	TCAAGCGGTC	AATTATCAAA	CGCGTTGCAA	CAACGGTAAC
6351	${\tt CTACAGGCAT}$	${\tt CGTGGTGTCA}$	${\tt CGCTCGTCGT}$	${\tt TTGGTATGGC}$	TTCATTCAGC
	GATGTCCGTA	GCACCACAGT	GCGAGCAGCA	AACCATACCG	AAGTAAGTCG
6401	${\tt TCCGGTTCCC}$	AACGATCAAG	${\tt GCGAGTTACA}$	TGATCCCCCA	TGTTGTGCAA
	AGGCCAAGGG	TTGCTAGTTC	CGCTCAATGT	ACTAGGGGGT	ACAACACGTT
6451	AAAAGCGGTT	AGCTCCTTCG	GTCCTCCGAT	CGTTGTCAGA	AGTAAGTTGG
	TTTTCGCCAA	TCGAGGAAGC	CAGGAGGCTA	GCAACAGTCT	TCATTCAACC
6501			GTTATGGCAG		
	GGCGTCACAA	TAGTGAGTAC	CAATACCGTC	GTGACGTATT	AAGAGAATGA
6551			CTTTTCTGTG		
	CAGTACGGTA	GGCATTCTAC	GAAAAGACAC	TGACCACTCA	TGAGTTGGTT
6601			TGCGGCGACC		
			ACGCCGCTGG		
6651			CCACATAGCA		
			GGTGTATCGT		
6701			GCGAAAACTC		
			CGCTTTTGAG		
6751			CCACTCGTGC		
			GGTGAGCACG		
6801			TCTGGGTGAG		
			AGACCCACTC		
6851		+	GGCGACACGG		
			CCGCTGTGCC		
6901			GAAGCATTTA		
			CTTCGTAAAT		
6951			ATTTAGAAAA		
			TAAATCTTTT		TCCCCAAGGC
7001			GCCACCTGAC		
	GCGTGTAAAG	GGGCTTTTCA	CGGTGGACTG	CAG	

# FIG. 10 (1-12)

### pCEP4W/hepEK

1	TCGAGCGGCC GCTTTAAACT CAATGGTGAT GGTGATGATG ACCGGTACGC
	AGCTCGCCGG CGAAATTTGA GTTACCACTA CCACTACTAC TGGCCATGCG
-3	H H H H H G T R T
51	GTAGAATCGA GACCGAGGAG AGGGTTAGGG ATAGGCTTAC CGAATTCGAG
	CATCTTAGCT CTGGCTCCTC TCCCAATCCC TATCCGAATG GCTTAAGCTC
-3	·TSDLGLLPNP!PKGFEL
101	CTGGGTCACC ATGCCGCTGG CTTCGGAGTG AGTCTTTATG GCCTGGAAGA
	GACCCAGTGG TACGGCGACC GAAGCCTCAC TCAGAAATAC CGGACCTTCT
-3	Q T V M G S A E S H T K I A Q F I
151	TCCACTCCCG GAAGTCACTG ACTTTGGTGT AGACGCCTGG CTTCTGGGCC
	AGGTGAGGGC CTTCAGTGAC TGAAACCACA TCTGCGGACC GAAGACCCGG
-3	·IWERFDSVKTYVGPKQAL
201	AGGGCACAGC CAGTGCCCCA ACTCACAATG CCACACAGCC GCCAACGTGG
	TCCCGTGTCG GTCACGGGGT TGAGTGTTAC GGTGTGTCGG CGGTTGCACC
-3	·LACGTGWSVIGCLRWRP
251	CGTCCGAGAG ATGCTGTCCT CACACACAAA GGGACCACCG CTGTCGCCCT
	GCAGGCTCTC TACGACAGGA GTGTGTGTTT CCCTGGTGGC GACAGCGGGA
-3	TRSISDECVFPGGSDGQ
301	GGCAGGCATC AATGCCACCC TCGGGGTAGC CAGCACAGAA CATCTTGGGC
	CCGTCCGTAG TTACGGTGGG AGCCCCATCG GTCGTGTCTT GTAGAACCCG
-3	·QCADIGGEPYGACFMKPK
351	TTGATCTGGT TTCCATAGAA ATCAGCGCCA TTGCAGACAT CATTGCTGAT
	AACTAGACCA AAGGTATCTT TAGTCGCGGT AACGTCTGTA GTAACGACTA
-3	·KIQNGYFDAGNCVDNSI
401	TATGGGGACT CGAGCCTCCT GGAGTACCCC GGCCTGTTGG CCATAGTACT
	ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA
-3	IPVRAEQLVGAQQGYYQ.
451	GCGTGTTGCC CCAGCCCGTC ACGGTACAGA TCTTGCCATC CACCAGGGCC
	CGCACAACGG GGTCGGGCAG TGCCATGTCT AGAACGGTAG GTGGTCCCGG
-3	QTNG WGT V TC I KG D V L A Q
501	TGGCCGGCAG CTGGGAGGCA CACAGGCTGG ATGTATTCTG TGAGGGGCAG
	ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC
-3	ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC Q G A A P L C V P Q I Y E T L P L
-3 551	ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC Q G A A P L C V P Q I Y E T L P L GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT
551	ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC  Q G A A P L C V P Q I Y E T L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA
551 -3	ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC  O G A A P L C V P Q I Y E T L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA  P S S L H V L A I D N S N E E S N
551	ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC  O G A A P L C V P Q I Y E T L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA  P S S L H V L A I D N S N E E S N  TGGGGTCCCG AAAGGGAAGA TAGCCCCCGT GGTAGACCAC AGCCTGCACC
551 -3 601	ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC  Q G A A P L C V P Q I Y E T L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA  P S S L H V L A I D N S N E E S N  TGGGGTCCCG AAAGGGAAGA TAGCCCCCGT GGTAGACCAC AGCCTGCACC  ACCCCAGGGC TTTCCCTTCT ATCGGGGGCA CCATCTGGTG TCGGACGTGG
-3 601	ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC  Q G A A P L C V P Q I Y E T L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA  P S S L H V L A I D N S N E E S N  TGGGGTCCCG AAAGGGAAGA TAGCCCCCGT GGTAGACCAC AGCCTGCACC  ACCCCAGGGC TTTCCCTTCT ATCGGGGGCA CCATCTGGTG TCGGACGTGG  N P D R F P L Y G G H Y V V A Q V G
551 -3 601	ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC  Q G A A P L C V P Q I Y E T L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA  P S S L H V L A I D N S N E E S N  TGGGGTCCCG AAAGGGAAGA TAGCCCCCGT GGTAGACCAC AGCCTGCACC  ACCCCAGGGC TTTCCCTTCT ATCGGGGGCA CCATCTGGTG TCGGACGTGG  N P D R F P L Y G G H Y V V A Q V G  CCCAGCTGCA GACCGTGGGG AGAGGCCTGG GCCACGGCAC CGGCAAACAC
-3 601 -3 651	ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC  Q G A A P L C V P Q I Y E T L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA  P S S L H V L A I D N S N E E S N  TGGGGTCCCG AAAGGGAAGA TAGCCCCCGT GGTAGACCAC AGCCTGCACC  ACCCCAGGGC TTTCCCTTCT ATCGGGGGCA CCATCTGGTG TCGGACGTGG  N P D R F P L Y G G H Y V V A Q V G  CCCAGCTGCA GACCGTGGGG AGAGGCCTGG GCCACGGCAC CGGCAAACAC  GGGTCGACGT CTGGCACCC TCTCCGGACC CGGTGCCGTG GCCGTTGTG
-3 601 -3 651	ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC  Q G A A P L C V P Q I Y E T L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA  P S S L H V L A I D N S N E E S N  TGGGGTCCCG AAAGGGAAGA TAGCCCCCGT GGTAGACCAC AGCCTGCACC  ACCCCAGGGC TTTCCCTTCT ATCGGGGGCA CCATCTGGTG TCGGACGTGG  N P D R F P L Y G G H Y V V A Q V G  CCCAGCTGCA GACCGTGGGG AGAGGCCTGG GCCACGGCAC CGGCAAACAC  GGGTCGACGT CTGGCACCC TCTCCGGACC CGGTGCCGTG GCCGTTGTG  G L Q L G H P S A Q A V A G A F V
-3 601 -3 651	ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC  Q G A A P L C V P Q I Y E T L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA  P S S L H V L A I D N S N E E S N  TGGGGTCCCG AAAGGGAAGA TAGCCCCCGT GGTAGACCAC AGCCTGCACC  ACCCCAGGGC TTTCCCTTCT ATCGGGGGCA CCATCTGGTG TCGGACGTGG  N P D R F P L Y G G H Y V V A Q V G  CCCAGCTGCA GACCGTGGGG AGAGGCCTGG GCCACGGCAC CGGCAAACAC  GGGTCGACGT CTGGCACCC TCTCCGGACC CGGTGCCGTG GCCGTTTGTG  G L Q L G H P S A Q A V A G A F V  TCGCCATCGG GACAGGACCC GGTTCCGCTC CGGGAAGCAG TGGGCGGCTG
-3 601 -3 651	ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC  Q G A A P L C V P Q I Y E T L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA  P S S L H V L A I D N S N E E S N  TGGGGTCCCG AAAGGGAAGA TAGCCCCCGT GGTAGACCAC AGCCTGCACC  ACCCCAGGGC TTTCCCTTCT ATCGGGGGCA CCATCTGGTG TCGGACGTGG  N P D R F P L Y G G H Y V V A Q V G  CCCAGCTGCA GACCGTGGGG AGAGGCCTGG GCCACGGCAC CGGCAAACAC  GGGTCGACGT CTGGCACCC TCTCCGGACC CGGTGCCGTG GCCGTTGTG  G L Q L G H P S A Q A V A G A F V

## FIGURE10-1

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751 TCAGCACCCA GTCCCCGGAG AGCAGGGATC CCCCACAGAG GTGTGCTCCA AGTCGTGGGT CAGGGGCCTC TCGTCCCTAG GGGGTGTCTC CACACGAGGT -3 ·TLV W D G S L L S G G C L H A G D TCATAGCGAA GGCTGACTTG CCACGGCCAC CGGCCCAAGC TGGTGTCCCG AGTATCGCTT CCGACTGAAC GGTGCCGGTG GCCGGGTTCG ACCACAGGGC ·DYR L S V Q W P W R G L 851 GCCTCCCACG ATCTTGTCGT CGTCGTCCAC GGGCAGCTTC CTGCGGCCAC CGGAGGGTGC TAGAACAGCA GCAGCAGGTG CCCGTCGAAG GACGCCGGTG G G V I K D D D D V P L K R R G C. 901 AGTCTTGGCA GATGGCGGCC AAGAAACGGC CTCTGGGGCA ATCACACACG TCAGAACCGT CTACCGCCGG TTCTTTGCCG GAGACCCCGT TAGTGTGTGC -3 ·CDQCIAALFRGRPCDCVS GAGATGACCT CCAGCAGCCT CTGGGTGTGG GGCAGCCTCC CCTCGTCCAC 951 CTCTACTGGA GGTCGTCGGA GACCCACACC CCGTCGGAGG GGAGCAGGTG SIVELLRQTHPLRGEDV ACAGAAGAAG CCCGACGTGC CATTGGCGCC CGCCGTTCGC ACGTCCAGCT 1001 TGTCTTCTTC GGGCTGCACG GTAACCGCGG GCGGCAAGCG TGCAGGTCGA S T G NAGATR CGGAGTGGGT CAGTGCCCTG AGGAAGCCCA TCTCCTCGCA GCTGAGTCCG 1051 GCCTCACCCA GTCACGGGAC TCCTTCGGGT AGAGGAGCGT CGACTCAGGC -3 ·ESHTLARL FGMEEC SLGA 1101 GCTACCCTGG CGTTAGAGCG CGAGGAGCAC AGCAGCCGCC ACGTCCCTTC CGATGGGACC GCAATCTCGC GCTCCTCGTG TCGTCGGCGG TGCAGGGAAG ·AVRANSR SSCLLRW CGTCTTGTCA AAGACCATGA GCCGAGCGTC CGCAGAGCTG ACCTGCACTG 1151 GCAGAACAGT TTCTGGTACT CGGCTCGCAG GCGTCTCGAC TGGACGTGAC T K D F V M L R A D A S S V GGTACAGCGG CTCCTGGTCA CTCCTATCCG GAGCGTCACC AGTGGAACCT 1201 CCATGTCGCC GAGGACCAGT GAGGATAGGC CTCGCAGTGG TCACCTTGGA EQDSRDP 1251 GGAACCCAGA GCAGCAGTAC CCATAGCAGG AGTGTGTCTG TCTCCATGGT CCTTGGGTCT CGTCGTCATG GGTATCGTCC TCACAGAGC AGAGGTACCA PVWLLLVWLLLTDTEM GGCGATCTGG TACCCAGCTT CTAGAGATCT GACGGTTCAC TAAACGAGCT 1301 CCGCTAGACC ATGGGTCGAA GATCTCTAGA CTGCCAAGTG ATTTGCTCGA CTGCTTATAT AGACCTCCCA CCGTACACGC CTACCGCCCA TTTGCGTCAA 1351 GACGAATATA TCTGGAGGGT GGCATGTGCG GATGGCGGGT AAACGCAGTT 1401 CGGGGCGGG TTATTACGAC ATTTTGGAAA GTCCCGTTGA TTTTGGTGCC GCCCCGCCC AATAATGCTG TAAAACCTTT CAGGGCAACT AAAACCACGG AAAACAAACT CCCATTGACG TCAATGGGGT GGAGACTTGG AAATCCCCGT 1451 TTTTGTTTGA GGGTAACTGC AGTTACCCCA CCTCTGAACC TTTAGGGGCA GAGTCAAACC GCTATCCACG CCCATTGGTG TACTGCCAAA ACCGCATCAC 1501 CTCAGTTTGG CGATAGGTGC GGGTAACCAC ATGACGGTTT TGGCGTAGTG 1551 CATGGTAATA GCGATGACTA ATACGTAGAT GTACTGCCAA GTAGGAAAGT GTACCATTAT CGCTACTGAT TATGCATCTA CATGACGGTT CATCCTTTCA

1601		CATGTACTGG GTACATGACC			
1651					
1651		TAGGGGGCGG ATCCCCCGCC			
1701	CAAGTGGGCA	GTTTACCGTA	AATACTCCAC	CCATTGACGT	CAATGGAAAG
	GTTCACCCGT	CAAATGGCAT	TTATGAGGTG	GGTAACTGCA	GTTACCTTTC
1751	тесстаттес	CGTTACTATG	GGAACATACG	тсаттаттса	CGTCAATGGG
1,31		GCAATGATAC			
1801	CGGGGGTCGT	TGGGCGGTCA	GCCAGGCGGG	CCATTTACCG	TAAGTTATGT
	GCCCCAGCA	ACCCGCCAGT	CGGTCCGCCC	GGTAAATGGC	ATTCAATACA
1851	AACGCGGAAC	TCCATATATG	GGCTATGAAC	TAATGACCCC	GTAATTGATT
	TTGCGCCTTG	AGGTATATAC	CCGATACTTG	ATTACTGGGG	CATTAACTAA
1901	АСТАТТААТА	ACTAGTCAAT	AATCAATGTC	AACATGGCGG	TCATATTGGA
		TGATCAGTTA			
1951	CATGAGCCAA	TATAAATGTA	САТАТТАТСА	ТАТАСАТАСА	ACGTATGCAA
1331	_	ATATTTACAT			
2001	TGGCCAATAG	CCAATATTGA	TTTATGCTAT	ATAACCAATG	ACTAATATGG
	ACCGGTTATC	GGTTATAACT	AAATACGATA	TATTGGTTAC	TGATTATACC
2051	CTAATTGCCA	ATATTGATTC	AATGTATAGA	TCTTCCATAC	CTACCAGTTC
	GATTAACGGT	TATAACTAAG	TTACATATCT	AGAAGGTATG	GATGGTCAAG
2101	TGCGCCTGCA	GCAATGCAAC	AACGTTGCCC	GGATCTGCGA	TGATAAGCTG
	ACGCGGACGT	CGTTACGTTG	TTGCAACGGG	CCTAGACGCT	ACTATTCGAC
2151	TCAAACATGA	GAATTGGTCG	ACTAGCTTGG	CACGCCAGAA	ATCCGCGCGG
	AGTTTGTACT	CTTAACCAGC	TGATCGAACC	GTGCGGTCTT	TAGGCGCGCC
2201	TGGTTTTTGG	GGGTCGGGG	TGTTTGGCAG	CCACAGACGC	CCGGTGTTCG
		CCCAGCCCCC			-
2251	тстсссссса	GTACATGCGG	тссатессса	GGCCATCCAA	AAACCATGGG
2232		CATGTACGCC			
2301		CAGTCCAGTC			
2501					GCGGGTTTTA
2351					CCCGTCCCTA
2331			_		GGGCAGGGAT
2401					ACGTTGGCTG
2401		GGTCACCGAT			
2451					
2451					GGAAAAGGAA CCTTTTCCTT
2501					
2501					TATCGACAGA ATAGCTGTCT
0551					
2551					GACCCAACAC CTGGGTTGTG
2601					
2601					GGGTTCCTTC
2651					
2651					CACCCCAMA
0701					GAGGGGATAA
2701					' ATCGGCGAGT
	GGAAACGGGA	GCCTGCTCAC	GACCCCGCAG	CCAAAGGTGA	TAGCCGCTCA

2751				GCGCTTCTGC CGCGAAGACG	
2001				GACGATTGCG	
2801				CTGCTAACGC	
2851	CCTGCGCCCA	AGCTGCATCA	TCGAAATTGC	CGTCAACCAA	GCTCTGATAG
	GGACGCGGGT	TCGACGTAGT	AGCTTTAACG	GCAGTTGGTT	CGAGACTATC
2901	AGTTGGTCAA	GACCAATGCG	GAGCATATAC	GCCCGGAGCC	GCGGCGATCC
	TCAACCAGTT	CTGGTTACGC	CTCGTATATG	CGGGCCTCGG	CGCCGCTAGG
2951	ТССААССТСС	GGATGCCTCC	GCTCGAAGTA	GCGCGTCTGC	TGCTCCATAC
				CGCGCAGACG	
3001	AAGCCAACCA	CGGCCTCCAG	AAGAAGATGT	TGGCGACCTC	GTATTGGGAA
3001				ACCGCTGGAG	
3051				ACCGCTGTTA	
3031				TGGCGACAAT	
3101				CGCGTGCACG	
2101				GCGCACGTGC	
3151				GCTCATCGAG	
2131				CGAGTAGCTC	
2201				GTTTGCCAGT	
3201				CAAACGGTCA	
2251				CCATGTAGTG	
3251				GGTACATCAC	
2201					
3301				TCGTCTGGCT AGCAGACCGA	
2254					
3351	• • • • • • • • • • • • • • • • • • • •			GGCTGCAGAA CCGACGTCTT	
3401				ACCCTGTGCA TGGGACACGT	
3451					CACTTCCGGA
				GTTACAGTTC	
3501					GATCTTTGTA
					CTAGAAACAT
3551					CGCCCTCCTA
		_			GCGGGAGGAT
3601					CTGCATCAGG
					GACGTAGTCC
3651					CAGACGTCGC
					GTCTGCAGCG
3701					GCGGCACGCT
					CGCCGTGCGA
3751					TTCGAGGCCA
	CAACTGCGAC	: AATTCGCCCA	GCGACGTCCC	C AGCGAGCCAC	AAGCTCCGGT
3801					CCGCCCCGAC
	GTGCGCAGT	GAATTATACO	G CTTCACCTGO	ACCCTGGCGC	GGCGGGGCTG
3851					GCGGGGTTTG
	ACGTAGACGO	C ACAAGCTTAA	A GCGGTTACTO	G TTCTGCGACO	CGCCCCAAAC

3901		GAACTAAAGA CTTGATTTCT			
3951		CGAGCAACGG GCTCGTTGCC			
4001		GCTACGTCTT CGATGCAGAA			
4051		ATGATTCTTC TACTAAGAAG			
4101		GCTGTCCAGG CGACAGGTCC			
4151		TCGCGGCTCT AGCGCCGAGA			
4201		GCGATTTATG CGCTAAATAC			
4251		AGGCGCCGCC TCCGCGGCGG			
4301		GGAGCCGGGC CCTCGGCCCG			
4351		GATTCACCAC CTAAGTGGTG			
4401		AATGCGCAAA TTACGCGTTT			
4451	GGCGGTAGAG	GTCGTCGGCG	TGCGCCGCGT	CGTTTTCCGG	
4501	TTTTCCGGCG	GTTGCTGGCG CAACGACCGC	AAAAAGGTAT	CCGAGGCGGG	GGGACTGCTC
4551					CGACAGGACT GCTGTCCTGA
4601	TATTTCTATG	GTCCGCAAAG	GGGGACCTTC	GAGGGAGCAC	CGCTCTCCTG
4651	AAGGCTGGGA	CGGCGAATGG	CCTATGGACA	GGCGGAAAGA	CCCTTCGGGA GGGAAGCCCT
4701	TCGCACCGCG	AAAGAGTATC	GAGTGCGACA	TCCATAGAGI	GTTCGGTGTA CAAGCCACAT
4751	CCAGCAAGCG	AGGTTCGACC	CGACACACGI	GCTTGGGGGG	GTTCAGCCCG GCAAGTCGGGC
4801	TGGCGACGCG	GAATAGGCCA	TTGATAGCAG	AACTCAGGTT	CCCGGTAAGA GGGCCATTCT
4851	GTGCTGAATA	A GCGGTGACCG	TCGTCGGTGA	CCATTGTCCT	A TTAGCAGAGC AATCGTCTCG
4901	CTCCATACAT	CCGCCACGAT	GTCTCAAGA	A CTTCACCACO	G CCTAACTACG C GGATTGATGC
4951	CGATGTGAT	TTCCTGTCAT	T AAACCATAGA	A CGCGAGACGA	GAAGCCAGTT CTTCGGTCAA
5001					AAACCACCGC TTTGGTGGCG

5051		GGTTTTTTTG CCAAAAAAAC			
5101		AGAAGATCCT			
3101		TCTTCTAGGA			
5151	TGGAACGAAA	ACTCACGTTA	AGGGATTTTG	GTCATGAGAT	TATCAAAAAG
	ACCTTGCTTT	TGAGTGCAAT	TCCCTAAAAC	CAGTACTCTA	ATAGTTTTTC
5201		TAGATCCTTT ATCTAGGAAA			
5251	AAAGTATATA	TGAGTAAACT	TGGTCTGACA	GTTACCAATG	CTTAATCAGT
	TTTCATATAT	ACTCATTTGA	ACCAGACTGT	CAATGGTTAC	GAATTAGTCA
5301	GAGGCACCTA	TCTCAGCGAT	CTGTCTATTT	CGTTCATCCA	TAGTTGCCTG
	CTCCGTGGAT	AGAGTCGCTA	GACAGATAAA	GCAAGTAGGT	ATCAACGGAC
5351	ACTCCCCGTC	GTGTAGATAA	CTACGATACG	GGAGGGCTTA	CCATCTGGCC
	TGAGGGGCAG	CACATCTATT	GATGCTATGC	CCTCCCGAAT	GGTAGACCGG
5401	CCAGTGCTGC	AATGATACCG	CGAGACCCAC	GCTCACCGGC	TCCAGATTTA
		TTACTATGGC			
5451	TCAGCAATAA	ACCAGCCAGC	CGGAAGGCC	GAGCGCAGAA	GTGGTCCTGC
	AGTCGTTATT	TGGTCGGTCG	GCCTTCCCGG	CTCGCGTCTT	CACCAGGACG
5501	AACTTTATCC	GCCTCCATCC	AGTCTATTAA	TTGTTGCCGG	GAAGCTAGAG
		CGGAGGTAGG			
5551	TAAGTAGTTC	GCCAGTTAAT	AGTTTGCGCA	ACGTTGTTGC	CATTGCTGCA
	ATTCATCAAG	CGGTCAATTA	TCAAACGCGT	TGCAACAACG	GTAACGACGT
5601	GGCATCGTGG	TGTCACGCTC	GTCGTTTGGT	ATGGCTTCAT	TCAGCTCCGG
	CCGTAGCACC	ACAGTGCGAG	CAGCAAACCA	TACCGAAGTA	AGTCGAGGCC
5651	TTCCCAACGA	TCAAGGCGAG	TTACATGATC	CCCCATGTTG	TGCAAAAAAG
	AAGGGTTGCT	AGTTCCGCTC	AATGTACTAG	GGGGTACAAC	ACGTTTTTTC
5701	CGGTTAGCTC	CTTCGGTCCT	CCGATCGTTG	TCAGAAGTAA	GTTGGCCGCA
	GCCAATCGAG	GAAGCCAGGA	GGCTAGCAAC	AGTCTTCATT	CAACCGGCGT
5751	GTGTTATCAC	TCATGGTTAT	GGCAGCACTG	CATAATTCTC	TTACTGTCAT
	CACAATAGTG	AGTACCAATA	CCGTCGTGAC	GTATTAAGAG	AATGACAGTA
5801	GCCATCCGTA	AGATGCTTTT	CTGTGACTGG	TGAGTACTCA	ACCAAGTCAT
	CGGTAGGCAT	TCTACGAAAA	GACACTGACC	ACTCATGAGT	TGGTTCAGTA
5851	TCTGAGAATA	GTGTATGCGG	CGACCGAGTT	GCTCTTGCCC	GGCGTCAACA
	AGACTCTTAT	CACATACGCC	GCTGGCTCAA	CGAGAACGGG	CCGCAGTTGT
5901	CGGGATAATA	CCGCGCCACA	TAGCAGAACT	TTAAAAGTGC	TCATCATTGG
	GCCCTATTAT	GGCGCGGTGT	ATCGTCTTGA	AATTTTCACG	AGTAGTAACC
5951	AAAACGTTCT	TCGGGGCGAA	AACTCTCAAG	GATCTTACCG	CTGTTGAGAT
	TTTTGCAAGA	AGCCCCGCTT	TTGAGAGTTC	CTAGAATGGC	GACAACTCTA
6001	CCAGTTCGAT	GTAACCCACT	CGTGCACCCA	ACTGATCTTC	AGCATCTTTT
	GGTCAAGCTA	CATTGGGTGA	GCACGTGGGT	TGACTAGAAG	TCGTAGAAAA
6051	ACTTTCACCA	GCGTTTCTGG	GTGAGCAAAA	ACAGGAAGGC	AAAATGCCGC
	TGAAAGTGGT	CGCAAAGACC	CACTCGTTTT	TGTCCTTCCG	TTTTACGGCG
6101	AAAAAAGGGA	ATAAGGGCGA	CACGGAAATG	TTGAATACTC	ATACTCTTCC
	TTTTTTCCCT	TATTCCCGCT	GTGCCTTTAC	AACTTATGAG	TATGAGAAGG
6151	TTTTTCAATA	TTATTGAAGC	ATTTATCAGG	GTTATTGTCT	CATGAGCGGA
	AAAAAGTTAT	AATAACTTCG	TAAATAGTCC	CAATAACAGA	GTACTCGCCT

6201			GAAAAATAAA		
	ATGTATAAAC	TTACATAAAT	CTTTTTATTT	GTTTATCCCC	AAGGCGCGTG
6251			CTGACGTCTA		
	TAAAGGGGCT	TTTCACGGTG	GACTGCAGAT	TCTTTGGTAA	TAATAGTACT
6301	CATTAACCTA	TAAAAATAGG	CGTATCACGA	GGCCCTTTCG	TCTTCAAGAA
	GTAATTGGAT	ATTTTTATCC	GCATAGTGCT	CCGGGAAAGC	AGAAGTTCTT
6351	TTCTCATGTT	TGACAGCTTA	TCATCGATAA	GCTGATCCTC	ACAGGCCGCA
	AAGAGTACAA	ACTGTCGAAT	AGTAGCTATT	CGACTAGGAG	TGTCCGGCGT
6401	CCCAGCTTTT	CTTCCGTTGC	CCCAGTAGCA	TCTCTGTCTG	GTGACCTTGA
	GGGTCGAAAA	GAAGGCAACG	GGGTCATCGT	AGAGACAGAC	CACTGGAACT
6451	AGAGGAAGAG	GAGGGGTCCC	GAGAATCCCC	ATCCCTACCG	TCCAGCAAAA
	TCTCCTTCTC	CTCCCCAGGG	CTCTTAGGGG	TAGGGATGGC	AGGTCGTTTT
6501	AGGGGGACGA	GGAATTTGAG	GCCTGGCTTG	AGGCTCAGGA	CGCAAATCTT
	TCCCCCTGCT	CCTTAAACTC	CGGACCGAAC	$\mathtt{TCCGAGTCCT}$	GCGTTTAGAA
6551	GAGGATGTTC	AGCGGGAGTT	TTCCGGGCTG	CGAGTAATTG	GTGATGAGGA
	CTCCTACAAG	TCGCCCTCAA	AAGGCCCGAC	GCTCATTAAC	CACTACTCCT
6601	CGAGGATGGT	TCGGAGGATG	GGGAATTTTC	AGACCTGGAT	CTGTCTGACA
	GCTCCTACCA	AGCCTCCTAC	CCCTTAAAAG	TCTGGACCTA	GACAGACTGT
6651	GCGACCATGA	AGGGGATGAG	GGTGGGGGG	CTGTTGGAGG	GGGCAGGAGT
	CGCTGGTACT	TCCCCTACTC	CCACCCCCC	GACAACCTCC	CCCGTCCTCA
6701	CTGCACTCCC	TGTATTCACT	GAGCGTCGTC	TAATAAAGAT	GTCTATTGAT
	GACGTGAGGG	ACATAAGTGA	CTCGCAGCAG	ATTATTTCTA	CAGATAACTA
6751	CTCTTTTAGT	GTGAATCATG	TCTGACGAGG	GGCCAGGTAC	AGGACCTGGA
	GAGAAAATCA	CACTTAGTAC	AGACTGCTCC	CCGGTCCATG	TCCTGGACCT
6801	AATGGCCTAG	GAGAGAAGGG	AGACACATCT	GGACCAGAAG	GCTCCGGCGG
	TTACCGGATC	CTCTCTTCCC	TCTGTGTAGA	CCTGGTCTTC	CGAGGCCGCC
6851	CAGTGGACCT	CAAAGAAGAG	GGGGTGATAA	CCATGGACGA	GGACGGGGAA
	GTCACCTGGA	GTTTCTTCTC	CCCCACTATT	GGTACCTGCT	CCTGCCCCTT
6901	GAGGACGAGG	ACGAGGAGGC	GGAAGACCAG	GAGCCCCGGG	CGGCTCAGGA
	CTCCTGCTCC	TGCTCCTCCG	CCTTCTGGTC	CTCGGGGCCC	GCCGAGTCCT
6951	TCAGGGCCAA	GACATAGAGA	TGGTGTCCGG	AGACCCCAAA	AACGTCCAAG
	AGTCCCGGTT	CTGTATCTCT	ACCACAGGCC	TCTGGGGTTT	TTGCAGGTTC
7001	TTGCATTGGC	TGCAAAGGGA	CCCACGGTGG	AACAGGAGCA	GGAGCAGGAG
	AACGTAACCG	ACGTTTCCCT	GGGTGCCACC	TTGTCCTCGT	CCTCGTCCTC
7051	CGGGAGGGC	AGGAGCAGGA	GGGGCAGGAG	CAGGAGGAGG	GGCAGGAGCA
	GCCCTCCCCG	TCCTCGTCCT	CCCCGTCCTC	GTCCTCCTCC	CCGTCCTCGT
7101	GGAGGAGGGG	CAGGAGGGC	AGGAGGGCA	GGAGGGGCAG	GAGCAGGAGG
	CCTCCTCCCC	GTCCTCCCCG	TCCTCCCCGT	CCTCCCCGTC	CTCGTCCTCC
7151	AGGGGCAGGA	GCAGGAGGAG	GGGCAGGAGG	GGCAGGAGGG	GCAGGAGCAG
	TCCCCGTCCT	CGTCCTCCTC	CCCGTCCTCC	CCGTCCTCCC	CGTCCTCGTC
7201	GAGGAGGGC	AGGAGCAGGA	GGAGGGGCAG	GAGGGGCAGG	AGCAGGAGGA
	CTCCTCCCCG	TCCTCGTCCT	CCTCCCCGTC	CTCCCCGTCC	TCGTCCTCCT
7251	GGGGCAGGAG	GGGCAGGAGG	GGCAGGAGCA	GGAGGAGGG	CAGGAGCAGG
	CCCCGTCCTC	CCCGTCCTCC	CCGTCCTCGT	CCTCCTCCCC	GTCCTCGTCC
7301	AGGAGGGCA	GGAGGGGCAG	GAGCAGGAGG	AGGGGCAGGA	GGGGCAGGAG
	TCCTCCCCGT	CCTCCCCGTC	CTCGTCCTCC	TCCCCGTCCT	CCCCGTCCTC

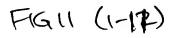
7351		AGGAGGAGGG TCCTCCTCCC			
7401		CAGGAGGGC			
	CCCCGTCCTC	GTCCTCCCCG	TCCTCGTCCT	CCTCCCCGTC	CTCCCCGTCC
7451		GCAGGAGGGG CGTCCTCCCC			
7501		AGGGGCAGGA			
	GTCCTCGTCC	TCCCCGTCCT	CCCCGTCCTC	GTCCTCCCCG	TCCTCCCCGT
7551		GGGCAGGAGG			
	CCTCGTCCTC	CCCGTCCTCC	CCGTCCTCGT	CCTCCTCCCC	GTCCTCCCCG
7601	AGGAGCAGGA	GGAGGGGCAG	GAGGGCAGG	AGCAGGAGGG	GCAGGAGGGG
	TCCTCGTCCT	CCTCCCCGTC	CTCCCCGTCC	TCGTCCTCCC	CGTCCTCCCC
7651		AGGGGCAGGA			
		TCCCCGTCCT			
7701		GAGGGCAGG			
		CTCCCCGTCC			
7751		GGAGGCAGTG			
		CCTCCGTCAC			
7801		AGGTAGTGGA			
		TCCATCACCT			
7851		GTCGTGAAAG			
	TCCCCCCCTT	CAGCACTTTC	TCGGTCCCCC	TCTCCACCAC	CTCCACCTCT
7901		AGGAGTCCCA			
	TTTCTCCGGG	TCCTCAGGGT	CATCAGTCAG	TACTACTACC	CCCACACCTCCAC
7951		CCCTCCAGGT			
,,,,,,		GGGAGGTCCA			
8001		TTGAATACCA			
0001	CGGCTAATAA	AACTTATGGT	CCAAGAAGGI	CCCCCTCTAC	GTGAGCCTGA
8051		GGAGCGATAG			
0001		CCTCGCTATC			
8101		TGGACCCCGG			
0101	CGGGTTCGTG	ACCTGGGGCC	CCACTCCCAC	TACCTCCCTC	GCGCAAAAA
8151		TTGGAAAGCA			
	CCTCCCACCA	AACCTTTCGT	AGCACCAGTT	CCTCCAACCT	ACCCGAAATT
8201		GCAGAAGGTT			
0202	ACTCTTGTAA	CGTCTTCCAA	ATTCTCCACA	CCTGGCTAGG	AGTCACGTAG
8251		CGACGAAGGA			
0231		GCTGCTTCCT			
8301		CCTCCCTTTA			
0501	CCATCATTCT	GGAGGGAAAT	GTTGGATTCC	CCTCCTTCA	CCCTTGCTAT
8351		CGTCTTACAC			
0331	AGGTGTTACA	GCAGAATGTG	CATTGAGTCG	ACACCCCTTT	GGAATGGCCC
8401					
0 <del>1</del> 0 T	GACCTCCCC	CCCACAACCT GGGTGTTGGA	CCCCCCARR	GGGAGTCCAT	TGTCTGTTAT
8451					
0471	AAGTACCACA	TTTTACAAAC	ACTATATATTT	GCTGAGGTTT	TGAAGGATGC
		AAAATGTTTG	AGIATAAA	CGACTCCAAA	ACTICCTACG

0504					
8501	GATTAAGGAC	CTTGTTATGA	CAAAGCCCGC	TCCTACCTGC	AATATCAGGG
		GAACAATACT			
8551		CAGCTTTGAC			
	ACTGACACAC	GTCGAAACTG	CTACCTCATC	TAAACGGAGG	GACCAAAGGT
8601	CCTATGGTGG	AAGGGGCTGC	CGCGGAGGGT	GATGACGGAG	ATGACGGAGA
	GGATACCACC	TTCCCCGACG	GCGCCTCCCA	CTACTGCCTC	TACTGCCTCT
8651		GATGGAGATG			
		CTACCTCTAC			
8701		CGCCCTCAAT			
		GCGGGAGTTA			
8751		TCCCCAGTAG			
		AGGGGTCATC			
8801		TGTCACCATT			
0001		ACAGTGGTAA			
8851					
0031		GTCACGTCAC			
0001		CAGTGCAGTG			
8901		AGCGACATTT			
0051		TCGCTGTAAA			
8951		CTCCTTAAAT			
0001		GAGGAATTTA			
9001		AAGCAGCGAA			
		TTCGTCGCTT			
9051		TAGCACTCCC			
		ATCGTGAGGG			
9101		AGGATAGCAT			
		TCCTATCGTA			
9151		GATATAGATT			
	TATGATGGGT	CTATATCTAA	TCCTATCGTA	${\tt TACGATGGGT}$	CTATATCTAA
9201		ATGCTACCCA			
	TCCTATCGGA	TACGATGGGT	CTATATTTAA	TCCTATCGTA	TATGATGGGT
9251	GATATAGATT	AGGATAGCAT	ATGCTACCCA	GATATAGATT	AGGATAGCCT
	CTATATCTAA	TCCTATCGTA	TACGATGGGT	CTATATCTAA	TCCTATCGGA
9301	ATGCTACCCA	GATATAGATT	AGGATAGCAT	ATGCTACCCA	GATATAGATT
	TACGATGGGT	CTATATCTAA	TCCTATCGTA	TACGATGGGT	CTATATCTAA
9351	AGGATAGCAT	ATGCTATCCA	GATATTTGGG	TAGTATATGC	TACCCAGATA
		TACGATAGGT			
9401		TAGCATATAC			
		ATCGTATATG			
9451		CAGATTAGGA			
		GTCTAATCCT			
9501		TACCCAGATA			
		ATGGGTCTAT			
9551		TAGCATATAC			
J J J L		ATCGTATATG			
9601					
7 O O T		TAGATTAGGA			
	OGGICIAI	ATCTAATCCT	AICGGATACG	AIGGGTCTAT	ATCTAATCCT

9651	TAGCATATGC	TATCCAGATA	TTTGGGTAGT	ATATGCTACC	CATGGCAACA
			AAACCCATCA		
9701			CGACCTCGTG		
			GCTGGAGCAC		
9751			AAGTGTGTGT		
			TTCACACACA		
9801			GCCCGCCCAC		
			CGGGCGGGTG		
9851			TGTGGGCAAG		
			ACACCCGTTC		
9901			${\tt GTTATTACAC}$		
			CAATAATGTG		
9951			GCTGACGCGT		
			CGACTGCGCA		
10001			TCTTTGTTTA		
40054			AGAAACAAAT		
10051			TGTTAGAGAC		
10101			ACAATCTCTG		
10101			CCCTTGTTAC		
10151			GGGAACAATG		
10151			GCCTGGGACA		
10001			CGGACCCTGT		
10201			TGTTGCCCAT		
10251			ACAACGGGTA		
10251	ACCECEACCE	GTCTTTACGG	CTTGTCCCCA	CCCCATGGAT	TTCTATTGTT
10301			GAACAGGGGT		
10301			ATTCCTACAC		
10351			TAAGGATGTG		
10331			TGTCATAGCA ACAGTATCGT		
10401					
10401			GGGCGTCACC CCCGCAGTGG		
10451			TCACAACTCA		
10431			AGTGTTGAGT		
10501			AGGCGAAGAT		
10301	TGCTTCCTCT	TACTTCTTCC	TCCGCTTCTA	ACTCCTCTCA	1 CACTGCCCG
10551			GCCCTTGTGA		
20002			CGGGAACACT		
10601			TAAATAAAAC		
	GCACCTTAGG	ACTGGGGTAC	ATTTATTTTG	GCACTGTCCA	CTACCCCACC
10651			GACCCTTTTA		
			CTGGGAAAAT		
10701			ATATGCTATT		
			TATACGATAA		
10751			AAGCATATGC		
	TCATATATGA	TGATGGGCCC	TTCGTATACG	ATGGGCAAAT	CCCAATTGTT

1080		TAAACACTAT ATTTGTGATA			
1085		CAGGCCCCTC			
1000		GTCCGGGGAG			
1090		CCTGGGAGGT			
2050		GGACCCTCCA			
1095	1 CAGCCAAGAG	TTACACATAA	AGGCAATGTT	GTGTTGCAGT	CCACAGACTG
		AATGTGTATT			
1100	)1 CAAAGTCTGC	TCCAGGATGA	AAGCCACTCA	GTGTTGGCAA	ATGTGCACAT
	GTTTCAGACG	AGGTCCTACT	TTCGGTGAGT	CACAACCGTT	TACACGTGTA
1105	1 CCATTTATAA	GGATGTCAAC	TACAGTCAGA	GAACCCCTTT	GTGTTTGGTC
	GGTAAATATI	CCTACAGTTG	ATGTCAGTCT	CTTGGGGAAA	CACAAACCAG
1110	1 CCCCCCGTG	TCACATGTGG	AACAGGGCCC	AGTTGGCAAG	TTGTACCAAC
	GGGGGGCAC	AGTGTACACC	TTGTCCCGGG	TCAACCGTTC	AACATGGTTG
1115	CAACTGAAGG	GATTACATGC	ACTGCCCCGC	GAAGAAGGGG	CAGAGATGCC
	GTTGACTTCC	CTAATGTACG	TGACGGGGCG	CTTCTTCCCC	GTCTCTACGG
1120	)1 GTAGTCAGGT	TTAGTTCGTC	CGGCGGCGGG	GCTCTAGAGT	CGACCGGTCA
	CATCAGTCCA	AATCAAGCAG	GCCGCCGCCC	CGAGATCTCA	GCTGGCCAGT
1125	TGGCTGCGCC	CCGACACCCG	CCAACACCCG	CTGACGCGCC	CTGACGGGCT
	ACCGACGCGG	GGCTGTGGGC	GGTTGTGGGC	GACTGCGCGG	GACTGCCCGA
1130	)1 TGTCTGCTCC	CGGCATCCGC	TTACAGACAA	GCTGTGACCG	TCTCCGGGAG
	ACAGACGAGG	GCCGTAGGCG	AATGTCTGTT	CGACACTGGC	AGAGGCCCTC
1135	51 CTGCATGTGT	CAGAGGTTTT	CACCGTCATC	ACCGAAACGC	GCGAGGCAGC
	GACGTACACA	GTCTCCAAAA	,GTGGCAGTAG	TGGCTTTGCG	CGCTCCGTCG
1140	)1 CGGATCATAA	TCAGCCATAC	CACATTTGTA	GAGGTTTTAC	TTGCTTTAAA
		AGTCGGTATG			
1145	51 AAACCTCCCC	ACCTCCCCT	GAACCTGAAA	CATAAAATGA	ATGCAATTGT
		TGGAGGGGGA			
1150		TTGTTTATTG			
		AACAAATAAC			
1155		TTTCACAAAT			
		' AAAGTGTTTA			
1160	)1 GGTTTGTCC <i>I</i>	AACTCATCAA	TGTATCTTAT	CATGTCTGGA	TCCCACGTGC
		TTGAGTAGTT			· · · · · · · · · · · · · · · · · · ·
1165		GCGGCCCAAA			
	TCCGCCCCTC	CGCCGGGTTT	CCCTCTAGGC	TGAGCAGACT	CCCGCTTCCG
1170	)1 GAAGACGCG	AAGAGGCCGC	AGAGCCGGCA	GCAGGCCGCG	GGAAGGAAGG
	CTTCTGCGCC	TTCTCCGGCG	TCTCGGCCGT	CGTCCGGCGC	CCTTCCTTCC
1175	TCCGCTGGAT	TGAGGGCCGA	AGGGACGTAG	CAGAAGGACG	TCCCGCGCAG
	AGGCGACCTA	ACTCCCGGCT	TCCCTGCATC	GTCTTCCTGC	AGGGCGCGTC
1180	01 AATCCAGGTO	GCAACACAGG	CGAGCAGCCA	AGGAAAGGAC	GATGATTTCC
•		CGTTGTGTCC			
1185		CACGGAATTG			
		GTGCCTTAAC			
1190		GGCAGGCGGC			
,		CCGTCCGCCG			

11951			GGAGCTGACA		
	CTTTCGCTTT	CAGGGCCTTT	CCTCGACTGT	CCACCACCGT	TACGGGGTTG
12001			AACACAGTGC		
	GTCACCCCCA	ACGCAGTCGT	TTGTGTCACG	TGTGGTGCGG	TGCAACGGAC
12051			ATAAAGAGAC		
	TGTTGCCCGG	TGTTGAGGAG	$\mathtt{TATTTCTCTG}$	${\tt TCGTTGGTCC}$	TAAATATGTT
12101			TACGGGAAGC		
	CCTCCTCTTT	TACTTTCGGT	${\tt ATGCCCTTCG}$	${\tt TTATCGTACT}$	ATGTTTCCGT
12151			${\tt AGCGTAAAAG}$		
	AATTTCGTCG	CATAGGTGTA	TCGCATTTTC	${\tt CTCGTTGTAT}$	CAATTCTTAT
12201	CCAGTCAATC	TTTCACAAAT	${\tt TTTGTAATCC}$	AGAGGTTGAT	TC
	GGTCAGTTAG	AAAGTGTTTA	AAACATTAGG	TCTCCAACTA	AG



1	TCGAGCGGCC GCTTTAAACT CAATGGTGAT GGTGATGATG ACCGGTACGC
	AGCTCGCCGG CGAAATTTGA GTTACCACTA CCACTACTAC TGGCCATGCG
-3	ннннн в т в т
51	GTAGAATCGA GACCGAGGAG AGGGTTAGGG ATAGGCTTAC CGAATTCGAG
	CATCTTAGCT CTGGCTCCTC TCCCAATCCC TATCCGAATG GCTTAAGCTC
-3	TSD L G L L P N P I P K G F E L
101	CTGGGTCACC ATGCCGCTGG CTTCGGAGTG AGTCTTTATG GCCTGGAAGA
	GACCCAGTGG TACGGCGACC GAAGCCTCAC TCAGAAATAC CGGACCTTCT
-3	Q T V M G S A E S H T K I A Q F I
151	TCCACTCCCG GAAGTCACTG ACTTTGGTGT AGACGCCTGG CTTCTGGGCC
	AGGTGAGGGC CTTCAGTGAC TGAAACCACA TCTGCGGACC GAAGACCCGG
-3	IWERFDS V KTY V G P K Q A L
201	AGGGCACAGC CAGTGCCCCA ACTCACAATG CCACACAGCC GCCAACGTGG
	TCCCGTGTCG GTCACGGGGT TGAGTGTTAC GGTGTGTCGG CGGTTGCACC
-3	·LACGTGWSVIGCLRWRP
251	CGTCCGAGAG ATGCTGTCCT CACACACAAA GGGACCACCG CTGTCGCCCT
	GCAGGCTCTC TACGACAGGA GTGTGTGTTT CCCTGGTGGC GACAGCGGGA
-3	TRSISDECVFPGGSDGQ
301	GGCAGGCATC AATGCCACCC TCGGGGTAGC CAGCACAGAA CATCTTGGGC
	CCGTCCGTAG TTACGGTGGG AGCCCCATCG GTCGTGTCTT GTAGAACCCG
-3	·QCADIGGEPYGACFMKPK
351	TTGATCTGGT TTCCATAGAA ATCAGCGCCA TTGCAGACAT CATTGCTGAT
	AACTAGACCA AAGGTATCTT TAGTCGCGGT AACGTCTGTA GTAACGACTA
	AACTAGACCA AAGGTATCTT TAGTCGCGGT AACGTCTGTA GTAACGACTA
-3	K ! Q N G Y F D A G N C V D N S !
-3 401	V. L. O. W. G. W.
	·K I Q N G Y F D A G N C V D N S I
	-K   Q N G Y F D A G N C V D N S   TATGGGGACT CGAGCCTCCT GGAGTACCCC GGCCTGTTGG CCATAGTACT
401	K I Q N G Y F D A G N C V D N S I  TATGGGGACT CGAGCCTCCT GGAGTACCCC GGCCTGTTGG CCATAGTACT  ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA
401	K I Q N G Y F D A G N C V D N S I  TATGGGGACT CGAGCCTCCT GGAGTACCCC GGCCTGTTGG CCATAGTACT  ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA  I P V R A E Q L V G A Q Q G Y Y Q
401	K I Q N G Y F D A G N C V D N S I  TATGGGGACT CGAGCCTCCT GGAGTACCCC GGCCTGTTGG CCATAGTACT  ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA  I P V R A E Q L V G A Q Q G Y Y Q  GCGTGTTGCC CCAGCCCGTC ACGGTACAGA TCTTGCCATC CACCAGGGCC
401 -3 451	-K ! Q N G Y F D A G N C V D N S !  TATGGGGACT CGAGCCTCCT GGAGTACCCC GGCCTGTTGG CCATAGTACT  ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA  I P V R A E Q L V G A Q Q G Y Y Q  GCGTGTTGCC CCAGCCCGTC ACGGTACAGA TCTTGCCATC CACCAGGGCC  CGCACAACGG GGTCGGGCAG TGCCATGTCT AGAACGGTAG GTGGTCCCGG
-3 451	K I Q N G Y F D A G N C V D N S I  TATGGGGACT CGAGCCTCCT GGAGTACCCC GGCCTGTTGG CCATAGTACT  ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA  I P V R A E Q L V G A Q Q G Y Y Q  GCGTGTTGCC CCAGCCCGTC ACGGTACAGA TCTTGCCATC CACCAGGGCC  CGCACAACGG GGTCGGGCAG TGCCATGTCT AGAACGGTAG GTGGTCCCGG  Q I N G W G I V I C I K G D V L A Q
-3 451	K I Q N G Y F D A G N C V D N S I  TATGGGGACT CGAGCCTCCT GGAGTACCCC GGCCTGTTGG CCATAGTACT  ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA  I P V R A E Q L V G A Q Q G Y Y Q  GCGTGTTGCC CCAGCCCGTC ACGGTACAGA TCTTGCCATC CACCAGGGCC  CGCACAACGG GGTCGGGCAG TGCCATGTCT AGAACGGTAG GTGGTCCCGG  Q I N G W G I V I C I K G D V L A Q  TGGCCGGCAG CTGGGAGGCA CACAGGCTGG ATGTATTCTG TGAGGGGCAG
401 -3 451 -3 501	K I Q N G Y F D A G N C V D N S I  TATGGGGACT CGAGCCTCCT GGAGTACCCC GGCCTGTTGG CCATAGTACT  ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA  I P V R A E Q L V G A Q Q G Y Y Q  GCGTGTTGCC CCAGCCCGTC ACGGTACAGA TCTTGCCATC CACCAGGGCC  CGCACAACGG GGTCGGGCAG TGCCATGTCT AGAACGGTAG GTGGTCCCGG  Q I N G W G I V I C I K G D V L A Q  TGGCCGGCAG CTGGGAGGCA CACAGGCTGG ATGTATTCTG TGAGGGGCAG  ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC
-3 451 -3 501	K I Q N G Y F D A G N C V D N S I  TATGGGGACT CGAGCCTCCT GGAGTACCCC GGCCTGTTGG CCATAGTACT  ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA  I P V R A E Q L V G A Q Q G Y Y Q GCGTGTTGCC CCAGCCCGTC ACGGTACAGA TCTTGCCATC CACCAGGGCC  CGCACAACGG GGTCGGGCAG TGCCATGTCT AGAACGGTAG GTGGTCCCGG  Q I N G W G I V I C I K G D V L A Q  TGGCCGGCAG CTGGGAGGCA CACAGGCTGG ATGTATTCTG TGAGGGGCAG  ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC  Q G A A P L C V P Q I Y E I L P L
-3 451 -3 501	K I Q N G Y F D A G N C V D N S I  TATGGGGACT CGAGCCTCCT GGAGTACCCC GGCCTGTTGG CCATAGTACT  ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA  I P V R A E Q L V G A Q Q G Y Y Q O  GCGTGTTGCC CCAGCCCGTC ACGGTACAGA TCTTGCCATC CACCAGGGCC  CGCACAACGG GGTCGGGCAG TGCCATGTCT AGAACGGTAG GTGGTCCCGG  Q I N G W G I V I C I K G D V L A Q  TGGCCGGCAG CTGGGAGGCA CACAGGCTGG ATGTATTCTG TGAGGGGCAG  ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC  Q G A A P L C V P Q I Y E I L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA  P S S L H V L A I D N S N E E S N
401 -3 451 -3 501 -3 551	K I Q N G Y F D A G N C V D N S I  TATGGGGACT CGAGCCTCCT GGAGTACCCC GGCCTGTTGG CCATAGTACT  ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA  I P V R A E Q L V G A Q Q G Y Y Q  GCGTGTTGCC CCAGCCCGTC ACGGTACAGA TCTTGCCATC CACCAGGGCC  CGCACAACGG GGTCGGGCAG TGCCATGTCT AGAACGGTAG GTGGTCCCGG  Q I N G W G I V I C I K G D V L A Q  TGGCCGGCAG CTGGGAGGCA CACAGGCTGG ATGTATTCTG TGAGGGGCAG  ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC  Q G A A P L C V P Q I Y E I L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA  P S S L H V L A I D N S N E E S N  TGGGGGTCCCG AAAGGGAAGA TAGCCCCCGT GGTAGACCAC AGCCTGCACC
401 -3 451 -3 501 -3 551	K I Q N G Y F D A G N C V D N S I  TATGGGGACT CGAGCCTCCT GGAGTACCCC GGCCTGTTGG CCATAGTACT  ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA  I P V R A E Q L V G A Q Q G Y Y Q O  GCGTGTTGCC CCAGCCCGTC ACGGTACAGA TCTTGCCATC CACCAGGGCC  CGCACAACGG GGTCGGGCAG TGCCATGTCT AGAACGGTAG GTGGTCCCGG  Q I N G W G I V I C I K G D V L A Q  TGGCCGGCAG CTGGGAGGCA CACAGGCTGG ATGTATTCTG TGAGGGGCAG  ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC  Q G A A P L C V P Q I Y E I L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA  P S S L H V L A I D N S N E E S N
401 -3 451 -3 501 -3 551	K I Q N G Y F D A G N C V D N S I  TATGGGGACT CGAGCCTCCT GGAGTACCCC GGCCTGTTGG CCATAGTACT  ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA  I P V R A E Q L V G A Q Q G Y Y Q  GCGTGTTGCC CCAGCCCGTC ACGGTACAGA TCTTGCCATC CACCAGGGCC  CGCACAACGG GGTCGGGCAG TGCCATGTCT AGAACGGTAG GTGGTCCCGG  Q I N G W G I V I C I K G D V L A Q  TGGCCGGCAG CTGGGAGGCA CACAGGCTGG ATGTATTCTG TGAGGGGCAG  ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC  Q G A A P L C V P Q I Y E I L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA  P S S L H V L A I D N S N E E S N  TGGGGTCCCG AAAGGGAAGA TAGCCCCCGT GGTAGACCAC AGCCTGCACC  ACCCCAGGGC TTTCCCTTCT ATCGGGGGCA CCATCTGGTG TCGGACGTGG  N P D R F P L Y G G H Y V V A Q V G
401 -3 451 -3 501 -3 551 -3 601	K I Q N G Y F D A G N C V D N S I  TATGGGGACT CGAGCCTCCT GGAGTACCCC GGCCTGTTGG CCATAGTACT  ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA  I P V R A E Q L V G A Q Q G Y Y Q Q G G GGGGGGGGGGGGGGGGG
401 -3 451 -3 501 -3 551 -3 601	K I Q N G Y F D A G N C V D N S I  TATGGGGACT CGAGCCTCCT GGAGTACCCC GGCCTGTTGG CCATAGTACT  ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA  I P V R A E Q L V G A Q Q G Y Y Q  GCGTGTTGCC CCAGCCCGTC ACGGTACAGA TCTTGCCATC CACCAGGGCC  CGCACAACGG GGTCGGGCAG TGCCATGTCT AGAACGGTAG GTGGTCCCGG  Q I N G W G I V I C I K G D V L A Q  TGGCCGGCAG CTGGGAGGCA CACAGGCTGG ATGTATTCTG TGAGGGGCAG  ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCCGTC  Q G A A P L C V P Q I Y E I L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA  P S S L H V L A I D N S N E E S N  TGGGGTCCCG AAAGGGAAGA TAGCCCCCGT GGTAGACCAC AGCCTGCACC  ACCCCAGGGC TTTCCCTTCT ATCGGGGGCA CCATCTGGTG TCGGACGTGG  N P D R F P L Y G G H Y V V A Q V G
401 -3 451 -3 501 -3 551 -3 601	K I Q N G Y F D A G N C V D N S I  TATGGGGACT CGAGCCTCT GGAGTACCCC GGCCTGTTGG CCATAGTACT  ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA  I P V R A E Q L V G A Q Q G Y Y Q O  GCGTGTTGCC CCAGCCCGTC ACGGTACAGA TCTTGCCATC CACCAGGGCC  CGCACAACGG GGTCGGGCAG TGCCATGTCT AGAACGGTAG GTGGTCCCGG  Q T N G W G T V T C I K G D V L A Q  TGGCCGGCAG CTGGGAGGCA CACAGGCTGG ATGTATTCTG TGAGGGGCAG  ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCGTC  Q G A A P L C V P Q I Y E T L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA  P S S L H V L A I D N S N E E S N  TGGGGTCCCG AAAGGGAAGA TAGCCCCCGT GGTAGACCAC AGCCTGCACC  ACCCCAGGGC TTTCCCTTCT ATCGGGGCA CCATCTGGTG TCGGACGTGG  N P D R F P L Y G G H Y V V V A Q V G  CCCAGCTGCA GACCGTGGG AGAGGCCTGG GCCACGGCAC CGGCAAACAC  GGGTCGACGT CTGGCACCC TCTCCGGACC CGGCACAGCACA
401 -3 451 -3 501 -3 551 -3 601 -3 651	K I Q N G Y F D A G N C V D N S I  TATGGGGACT CGAGCCTCCT GGAGTACCCC GGCCTGTTGG CCATAGTACT  ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA  I P V R A E Q L V G A Q Q G Y Y Q  GCGTGTTGCC CCAGCCCGTC ACGGTACAGA TCTTGCCATC CACCAGGGCC  CGCACAACGG GGTCGGCAG TGCCATGTCT AGAACGGTAG GTGGTCCCGG  Q T N G W G T V T C I K G D V L A Q  TGGCCGGCAG CTGGGAGGCA CACAGGCTGG ATGTATTCTG TGAGGGGCAG  ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCGTC  Q G A A P L C V P Q I Y E T L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA  P S S L H V L A I D N S N E E S N  TGGGGGTCCCG AAAGGGAAGA TAGCCCCCGT GGTAGACCAC AGCCTGCACC  ACCCCAGGGC TTTCCCTTCT ATCGGGGGCA CCATCTGGTG TCGGACGTGG  N P D R F P L Y G G H Y V V V A Q V G  CCCAGCTGCA GACCGTGGGG AGAGGCCTGG GCCACGGCAC CGGCAAACAC  GGGTCGACGT CTGGCACCC TCTCCGGACC CGGTGCCGTG GCCGTTGTG  G L Q L G H P S A Q A V A G A F V  TCGCCATCGG GACAGGACCC GGTTCCGCTC CGGGGAAGCAG TGGGCCGCTG
401 -3 451 -3 501 -3 651 -3	K I Q N G Y F D A G N C V D N S I  TATGGGGACT CGAGCCTCT GGAGTACCCC GGCCTGTTGG CCATAGTACT  ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA  I P V R A E Q L V G A Q Q G Y Y Q O  GCGTGTTGCC CCAGCCCGTC ACGGTACAGA TCTTGCCATC CACCAGGGCC  CGCACAACGG GGTCGGGCAG TGCCATGTCT AGAACGGTAG GTGGTCCCGG  Q T N G W G T V T C I K G D V L A Q  TGGCCGGCAG CTGGGAGGCA CACAGGCTGG ATGTATTCTG TGAGGGGCAG  ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCGTC  Q G A A P L C V P Q I Y E T L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA  P S S L H V L A I D N S N E E S N  TGGGGTCCCG AAAGGGAAGA TAGCCCCCGT GGTAGACCAC AGCCTGCACC  ACCCCAGGGC TTTCCCTTCT ATCGGGGCA CCATCTGGTG TCGGACGTGG  N P D R F P L Y G G H Y V V V A Q V G  CCCAGCTGCA GACCGTGGG AGAGGCCTGG GCCACGGCAC CGGCAAACAC  GGGTCGACGT CTGGCACCC TCTCCGGACC CGGCACAGCACA
401 -3 451 -3 501 -3 651 -3	K I Q N G Y F D A G N C V D N S I  TATGGGGACT CGAGCCTCCT GGAGTACCCC GGCCTGTTGG CCATAGTACT  ATACCCCTGA GCTCGGAGGA CCTCATGGGG CCGGACAACC GGTATCATGA  I P V R A E Q L V G A Q Q G Y Y Q  GCGTGTTGCC CCAGCCCGTC ACGGTACAGA TCTTGCCATC CACCAGGGCC  CGCACAACGG GGTCGGCAG TGCCATGTCT AGAACGGTAG GTGGTCCCGG  Q T N G W G T V T C I K G D V L A Q  TGGCCGGCAG CTGGGAGGCA CACAGGCTGG ATGTATTCTG TGAGGGGCAG  ACCGGCCGTC GACCCTCCGT GTGTCCGACC TACATAAGAC ACTCCCGTC  Q G A A P L C V P Q I Y E T L P L  GGGACTGGAG AGGTGGACCA GGGCAATATC GTTGCTGTTC TCCTCGCTGT  CCCTGACCTC TCCACCTGGT CCCGTTATAG CAACGACAAG AGGAGCGACA  P S S L H V L A I D N S N E E S N  TGGGGGTCCCG AAAGGGAAGA TAGCCCCCGT GGTAGACCAC AGCCTGCACC  ACCCCAGGGC TTTCCCTTCT ATCGGGGGCA CCATCTGGTG TCGGACGTGG  N P D R F P L Y G G H Y V V V A Q V G  CCCAGCTGCA GACCGTGGGG AGAGGCCTGG GCCACGGCAC CGGCAAACAC  GGGTCGACGT CTGGCACCC TCTCCGGACC CGGTGCCGTG GCCGTTGTG  G L Q L G H P S A Q A V A G A F V  TCGCCATCGG GACAGGACCC GGTTCCGCTC CGGGGAAGCAG TGGGCCGCTG

751	TCAGCACCCA GTCC				
	AGTCGTGGGT CAG	GGCCTC TC	GTCCCTAG	GGGGTGTCTC	CACACGAGGT
-3	·TLVWD	G S L	L S G	G C L	H A G D
801	TCATAGCGAA GGCT				
	AGTATCGCTT CCG	CTGAAC GO	TGCCGGTG	GCCGGGTTCG	ACCACAGGGC
-3	$\cdot  D  Y  R  L   S$	V Q W	/ P W F	R G L S	TDR
851	GCCTCCCACG ATC	TGTCGT CO	TCGTCCAC	GGGCAGCTTC	CTGCGGCCAC
	CGGAGGGTGC TAG	ACAGCA GO	AGCAGGTG	CCCGTCGAAG	GACGCCGGTG
-3	GGVI	C D D	D D V	PLK	R R G C
901	AGTCTTGGTC CGGA	AGCGTCA CO	CAGTGGAAC	CTGGAACCCA	GAGCAGCAGT
	TCAGAACCAG GCCT	CGCAGT GG	TCACCTTG	GACCTTGGGT	CTCGTCGTCA
-3	·CDQDP	A D G	T S G	P V W	L L L V
951	ACCCATAGCA GGAC	STGTGTC TO	TCTCCATG	GTGGCGATCT	GGTACCCAGC
	TGGGTATCGT CCTC	CACACAG AC	CAGAGGTAC	CACCGCTAGA	CCATGGGTCG
-3	·VWLLL	TDT	EM		
1001	TTCTAGAGAT CTG	ACGGTTC AC	CTAAACGAG	CTCTGCTTAT	ATAGACCTCC
	AAGATCTCTA GACT	GCCAAG TO	SATTTGCTC	GAGACGAATA	TATCTGGAGG
1051	CACCGTACAC GCCT	TACCGCC CA	ATTTGCGTC	AACGGGGCGG	GGTTATTACG
	GTGGCATGTG CGG	ATGGCGG GT	TAAACGCAG	TTGCCCCGCC	CCAATAATGC
1101	ACATTTTGGA AAGT	CCCGTT GA	ATTTTGGTG	ССААААСААА	CTCCCATTGA
	TGTAAAACCT TTC	AGGGCAA CI	TAAAACCAC	GGTTTTGTTT	GAGGGTAACT
1151	CGTCAATGGG GTGC	GAGACTT GO	SAAATCCCC	GTGAGTCAAA	CCGCTATCCA
	GCAGTTACCC CAC	CTCTGAA CO	TTTAGGGG	CACTCAGTTT	GGCGATAGGT
1201	CGCCCATTGG TGT	ACTGCCA AA	ACCGCATC	ACCATGGTAA	TAGCGATGAC
	GCGGGTAACC ACA	GACGGT TI	TTGGCGTAG	TGGTACCATT	ATCGCTACTG
1251	TAATACGTAG ATG	TACTGCC A	GTAGGAAA	GTCCCGTAAG	GTCATGTACT
	ATTATGCATC TACA	ATGACGG TO	CATCCTTT	CAGGGCATTC	CAGTACATGA
1301	GGGCATAATG CCAC	GCGGGC C	ATTTACCGT	CATTGACGTC	AATAGGGGGC
	CCCGTATTAC GGT	CCGCCCG G1	TAAATGGCA	GTAACTGCAG	TTATCCCCCG
1351	GGACTTGGCA TATO	GATACAC TI	GATGTACT	GCCAAGTGGG	CAGTTTACCG
	CCTGAACCGT ATAC	CTATGTG A	ACTACATGA	CGGTTCACCC	GTCAAATGGC
1401	TAAATACTCC ACC	CATTGAC G	CAATGGAA	AGTCCCTATT	GGCGTTACTA
	ATTTATGAGG TGGG	STAACTG C	AGTTACCTT	TCAGGGATAA	CCGCAATGAT
1451	TGGGAACATA CGT				
	ACCCTTGTAT GCA	GTAATAA CI	TGCAGTTAC	CCGCCCCCAG	CAACCCGCCA
1501	CAGCCAGGCG GGC				
	GTCGGTCCGC CCG	GTAAATG GO	CATTCAATA	CATTGCGCCT	TGAGGTATAT
1551	TGGGCTATGA ACT				
	ACCCGATACT TGA	TACTGG GO	GCATTAACT	AATGATAATT	ATTGATCAGT
1601	ATAATCAATG TCA				
	TATTAGTTAC AGT	rgtaccg co	CAGTATAAC	CTGTACTCGG	TTATATTTAC
1651	TACATATTAT GAT				
	ATGTATAATA CTA	TATCTAT G	TTGCATACG	TTACCGGTTA	TCGGTTATAA
1701	GATTTATGCT ATA				
	CTAAATACGA TAT	ATTGGTT AG	CTGATTATA	${\tt CCGATTAACG}$	GTTATAACTA

1751			ACCTACCAGT		
			TGGATGGTCA		
1801			GATGATAAGC		
			CTACTATTCG		
1851			AAATCCGCGC		
			TTTAGGCGCG		
1901			GCCCGGTGTT		
			CGGGCCACAA		
1951			AAAAACCATG		
			TTTTTGGTAC		
2001			AACGCCCAAA		
0054			TTGCGGGTTT		
2051			ACCCCGTCCC		
0101			TGGGGCAGGG		
2101			CGACGTTGGC		
21.51			GCTGCAACCG		
2151			GGGGAAAAGG		
2201			CCCCTTTTCC		
2201			GGTATCGACA CCATAGCTGT		
2251					
2231			ACGACCCAAC TGCTGGGTTG		
2301					
2301			GCGGGTTCCT CGCCCAAGGA	<del>-</del>	
2351			ATCTCCCCTA		
2331			TAGAGGGGAT		
2401			CTATCGGCGA		
2401			GATAGCCGCT		
2451			GCGGGCGATT		
2431			CGCCCGCTAA		
2501			CGTCGCATCG		
			GCAGCGTAGC		
2551			AAGCTCTGAT		
			TTCGAGACTA		
2601			CCGCGGCGAT		
	~				GGCCTACGGA
2651	CCGCTCGAAG	TAGCGCGTCT	GCTGCTCCAT	ACAAGCCAAC	CACGGCCTCC
					GTGCCGGAGG
2701					CATCGCCTCG
					GTAGCGGAGC
2751	CTCCAGTCAA	TGACCGCTGT	TATGCGGCCA	TTGTCCGTCA	GGACATTGTT
					CCTGTAACAA
2801	GGAGCCGAAA	TCCGCGTGCA	CGAGGTGCCG	GACTTCGGGG	CAGTCCTCGG
	CCTCGGCTTT	AGGCGCACGT	GCTCCACGGC	CTGAAGCCCC	GTCAGGAGCC
2851	CCCAAAGCAT	CAGCTCATCG	AGAGCCTGCG	CGACGGACGC	ACTGACGGTG
	GGGTTTCGTA	GTCGAGTAGC	TCTCGGACGC	GCTGCCTGCG	TGACTGCCAC

## FLGURE 11-3

2901		CAGTTTGCCA			
2051		GTCAAACGGT			
2951		CGCCATGTAG			
2001		GCGGTACATC			
3001		GCTCGTCTGG			
2054		CGAGCAGACC			
3051		CCGGCTGCAG			
2101		GGCCGACGTC			
3101		ACACCCTGTG			
2161		TGTGGGACAC			
3151		CCCAATGTCA			
2001		GGGTTACAGT			
3201		GATAAACATA CTATTTGTAT			
3251					
3231		AGGACATATC TCCTGTATAG			
3301		GCCCTCCGAG			
3301		CGGGAGGCTC			
3351		GAAACTTCTC			
3331		CTTTGAAGAG			
3401		TGCCCGGGAT			
3101		ACGGGCCCTA			
3451		GGTCGCTCGG			
		CCAGCGAGCC			
3501		CCTGGGACCG			
		GGACCCTGGC			
3551		ACAAGACGCT			
		TGTTCTGCGA			
3601		ТАТАТТТСТТ			
		ATATAAAGAA			
3651	GGGCCACGGG	GATGAAGCAG	GGCATGGCGG	CCGACGCGCT	GGGCTACGTC
		CTACTTCGTC			
3701	TTGCTGGCGT	TCGCGACGCG	AGGCTGGATG	GCCTTCCCCA	TTATGATTCT
	AACGACCGCA	AGCGCTGCGC	TCCGACCTAC	CGGAAGGGGT	AATACTAAGA
3751	TCTCGCTTCC	GGCGGCATCG	GGATGCCCGC	GTTGCAGGCC	ATGCTGTCCA
	AGAGCGAAGG	CCGCCGTAGC	CCTACGGGCG	CAACGTCCGG	TACGACAGGT
3801		TGACGACCAT			
	CCGTCCATCT	ACTGCTGGTA	GTCCCTGTCG	AAGTTCCTAG	CGAGCGCCGA
3851					CGGCGATTTA
		ATTGAAGCTA			
3901					GTAGGCGCCG
		CGCTCGTGTA			
3951		TGTCTGCCTC			
		ACAGACGGAG			
4001		CCTGAATGGA			
	CGGTGGAGCT	GGACTTACCT	TCGGCCGCCG	TGGAGCGATT	GCCTAAGTGG

4051		TTGGAGCCAA AACCTCGGTT			
4101					
4101		TGGCAGAACA ACCGTCTTGT			
4151	GCACGCGGCG	CAGCAAAAGG	CCAGGAACCG	TAAAAAGGCC	GCGTTGCTGG
	CGTGCGCCGC	GTCGTTTTCC	GGTCCTTGGC	ATTTTTCCGG	CGCAACGACC
4201		TAGGCTCCGC			
		ATCCGAGGCG			
4251	TCAAGTCAGA	GGTGGCGAAA	CCCGACAGGA	CTATAAAGAT	ACCAGGCGTT
	AGTTCAGTCT	CCACCGCTTT	GGGCTGTCCT	${\tt GATATTTCTA}$	TGGTCCGCAA
4301	TCCCCCTGGA	AGCTCCCTCG	TGCGCTCTCC	TGTTCCGACC	CTGCCGCTTA
	AGGGGGACCT	TCGAGGGAGC	ACGCGAGAGG	ACAAGGCTGG	GACGGCGAAT
4351	CCGGATACCT	GTCCGCCTTT	CTCCCTTCGG	GAAGCGTGGC	GCTTTCTCAT
	GGCCTATGGA	CAGGCGGAAA	GAGGGAAGCC	CTTCGCACCG	CGAAAGAGTA
4401	AGCTCACGCT	GTAGGTATCT	CAGTTCGGTG	TAGGTCGTTC	GCTCCAAGCT
		CATCCATAGA			
4451	GGGCTGTGTG	CACGAACCCC	CCGTTCAGCC	CGACCGCTGC	GCCTTATCCG
		GTGCTTGGGG			
4501		TCTTGAGTCC			
1001		AGAACTCAGG			
4551		CTGGTAACAG			·
1331		GACCATTGTC			
4601		TTGAAGTGGT			
.4001		AACTTCACCA			
4651		CTGCGCTCTG			
4031		GACGCGAGAC			
4701					
4/01		GATCCGGCAA	_		
4751		CTAGGCCGTT			
4751		CAGCAGATTA			
4001		GTCGTCTAAT			
4801		TTCTACGGGG			
		AAGATGCCCC			
4851		TGGTCATGAG			
					GGATCTAGGA
4901		AAATGAAGTT			
		TTTACTTCAA			
4951		CAGTTACCAA			
	GAACCAGACT	GTCAATGGTT	ACGAATTAGT	CACTCCGTGG	ATAGAGTCGC
5001		TTCGTTCATC			
	TAGACAGATA	AAGCAAGTAG	GTATCAACGG	ACTGAGGGGC	AGCACATCTA
5051		CGGGAGGGCT			
	TTGATGCTAT	GCCCTCCCGA	ATGGTAGACC	GGGGTCACGA	CGTTACTATG
5101	CGCGAGACCC	ACGCTCACCG	GCTCCAGATT	TATCAGCAAT	AAACCAGCCA
	GCGCTCTGGG	TGCGAGTGGC	CGAGGTCTAA	ATAGTCGTTA	TTTGGTCGGT
5151	GCCGGAAGGG	CCGAGCGCAG	AAGTGGTCCT	GCAACTTTAT	CCGCCTCCAT
	CGGCCTTCCC	GGCTCGCGTC	TTCACCAGGA	CGTTGAAATA	GGCGGAGGTA

5201	CCAGTCTATT	AATTGTTGCC	GGGAAGCTAG	AGTAAGTAGT	TCGCCAGTTA
5054		TTAACAACGG			
5251		CAACGTTGTT			
		GTTGCAACAA			
5301		GTATGGCTTC			
		CATACCGAAG			
5351		TCCCCCATGT			
		AGGGGGTACA			
5401		TGTCAGAAGT			
		ACAGTCTTCA			
5451		TGCATAATTC			
	TACCGTCGTG	ACGTATTAAG	AGAATGACAG	TACGGTAGGC	ATTCTACGAA
5501		GGTGAGTACT			
		CCACTCATGA			
5551	GGCGACCGAG	${\tt TTGCTCTTGC}$	CCGGCGTCAA	CACGGGATAA	TACCGCGCCA
	CCGCTGGCTC	AACGAGAACG	GGCCGCAGTT	${\tt GTGCCCTATT}$	ATGGCGCGGT
5601		CTTTAAAAGT			
	GTATCGTCTT	GAAATTTTCA	CGAGTAGTAA	${\tt CCTTTTGCAA}$	GAAGCCCCGC
5651		AGGATCTTAC			
	TTTTGAGAGT	TCCTAGAATG	GCGACAACTC	${\tt TAGGTCAAGC}$	TACATTGGGT
5701		CAACTGATCT			
		GTTGACTAGA			
5751		AAACAGGAAG			
	CCCACTCGTT	TTTGTCCTTC	CGTTTTACGG	${\tt CGTTTTTTCC}$	CTTATTCCCG
5801		TGTTGAATAC			
		ACAACTTATG			
5851		GGGTTATTGT			
		CCCAATAACA			
5901		AACAAATAGG			
		TTGTTTATCC			
5951		TAAGAAACCA			
		ATTCTTTGGT			
6001		GAGGCCCTTT			
		CTCCGGGAAA			
6051		AAGCTGATCC			
		TTCGACTAGG			
6101		CATCTCTGTC			
		GTAGAGACAG			
6151		CCATCCCTAC			
		GGTAGGGATG			
6201		TGAGGCTCAG			
		ACTCCGAGTC			
6251		TGCGAGTAAT			
		ACGCTCATTA			
6301		TCAGACCTGG			
	ACCCCTTAAA	AGTCTGGACC	TAGACAGACT	GTCGCTGGTA	CTTCCCCTAC

6351			GGGGGCAGGA		
C 4 0 1			CCCCCGTCCT		
6401			ATGTCTATTG		
			TACAGATAAC		
6451			ACAGGACCTG		
			TGTCCTGGAC		
6501			${\tt AGGCTCCGGC}$		
			TCCGAGGCCG		
6551			GAGGACGGGG		
	TCCCCCACTA	TTGGTACCTG	$\mathtt{CTCCTGCCCC}$	${\tt TTCTCCTGCT}$	CCTGCTCCTC
6601	GCGGAAGACC	AGGAGCCCCG	GGCGGCTCAG	GATCAGGGCC	AAGACATAGA
	CGCCTTCTGG	TCCTCGGGGC	CCGCCGAGTC	CTAGTCCCGG	TTCTGTATCT
6651	GATGGTGTCC	GGAGACCCCA	AAAACGTCCA	AGTTGCATTG	GCTGCAAAGG
	CTACCACAGG	CCTCTGGGGT	TTTTGCAGGT	TCAACGTAAC	CGACGTTTCC
6701	GACCCACGGT	GGAACAGGAG	CAGGAGCAGG	AGCGGGAGGG	GCAGGAGCAG
	CTGGGTGCCA	CCTTGTCCTC	GTCCTCGTCC	TCGCCCTCCC	CGTCCTCGTC
6751	GAGGGGCAGG	AGCAGGAGGA	GGGGCAGGAG	CAGGAGGAGG	GGCAGGAGGG
	CTCCCCGTCC	TCGTCCTCCT	CCCCGTCCTC	GTCCTCCTCC	CCGTCCTCCC
6801	GCAGGAGGGG	CAGGAGGGC	AGGAGCAGGA	GGAGGGGCAG	GAGCAGGAGG
			TCCTCGTCCT		
6851	AGGGGCAGGA	GGGGCAGGAG	GGGCAGGAGC	AGGAGGAGGG	GCAGGAGCAG
	TCCCCGTCCT	CCCCGTCCTC	CCCGTCCTCG	TCCTCCTCCC	CGTCCTCGTC
6901	GAGGAGGGC	AGGAGGGCA	GGAGCAGGAG	GAGGGCAGG	AGGGGCAGGA
			CCTCGTCCTC		
6951	GGGGCAGGAG	CAGGAGGAGG	GGCAGGAGCA	GGAGGAGGGG	CAGGAGGGC
			CCGTCCTCGT		
7001	AGGAGCAGGA	GGAGGGGCAG	GAGGGGCAGG	AGGGGCAGGA	GCAGGAGGAG
			CTCCCCGTCC		
7051	GGGCAGGAGC	AGGAGGGCA	GGAGGGGCAG	GAGGGCAGG	AGCAGGAGGG
			CCTCCCCGTC		
7101	GCAGGAGCAG	GAGGAGGGC	AGGAGGGGCA	GGAGGGGCAG	GAGCAGGAGG
	CGTCCTCGTC	CTCCTCCCG	TCCTCCCCGT	CCTCCCCGTC	CTCGTCCTCC
7151	GGCAGGAGCA	GGAGGGGCAG	GAGCAGGAGG	GGCAGGAGCA	GGAGGGGCAG
	CCGTCCTCGT	CCTCCCCGTC	CTCGTCCTCC	CCGTCCTCGT	CCTCCCCGTC
7201	GAGGGGCAGG	AGCAGGAGGG	GCAGGAGGG	CAGGAGCAGG	AGGGGCAGGA
	CTCCCCGTCC	TCGTCCTCCC	CGTCCTCCCC	GTCCTCGTCC	TCCCCGTCCT
7251	GGGGCAGGAG	CAGGAGGAGG	GGCAGGAGGG	GCAGGAGCAG	GAGGAGGGC
	CCCCGTCCTC	${\tt GTCCTCCTCC}$	CCGTCCTCCC	CGTCCTCGTC	CTCCTCCCG
7301	AGGAGGGGCA	GGAGCAGGAG	GGGCAGGAGG	GGCAGGAGCA	GGAGGGGCAG
	TCCTCCCCGT	CCTCGTCCTC	CCCGTCCTCC	CCGTCCTCGT	CCTCCCCGTC
7351	GAGGGCAGG	AGCAGGAGGG	GCAGGAGGGG	CAGGAGCAGG	AGGAGGGGCA
			CGTCCTCCC		
7401			AGGAGGTGGA		
			TCCTCCACCT		
7451	TGGAGGCCGG	GGTCGAGGAG	GTAGTGGAGG	CCGGGGTCGA	GGAGGTAGTG
			CATCACCTCC		

FIGURE 11-7

7501	GAGGCCGCCG CTCCGGCGGC	GGGTAGAGGA CCCATCTCCT	CGTGAAAGAG GCACTTTCTC	CCAGGGGGG	AAGTCGTGAA
7551					
,331	TCTCGGTCCC	CCTCTCCAGC	TGGACGTGGA ACCTGCACCT	CTTTTCTCCG	GGTCCTCAGG
7601	CAGTAGTCAG	TCATCATCAT	CCGGGTCTCC	ACCGCGCAGG	CCCCCTCCAG
	GTCATCAGTC	AGTAGTAGTA	GGCCCAGAGG	TGGCGCGTCC	GGGGGAGGTC
7651			CCTGTAGGGG		
	CATCTTCCGG	TAAAAAGGTG	GGACATCCCC	TTCGGCTAAT	AAAACTTATG
7701	CACCAAGAAG	GTGGCCCAGA	TGGTGAGCCT	GACGTGCCCC	CGGGAGCGAT
	GTGGTTCTTC	CACCGGGTCT	ACCACTCGGA	CTGCACGGGG	GCCCTCGCTA
7751	AGAGCAGGGC	CCCGCAGATG	ACCCAGGAGA	AGGCCCAAGC	ACTGGACCCC
	TCTCGTCCCG	GGGCGTCTAC	TGGGTCCTCT	TCCGGGTTCG	TGACCTGGGG
7801			AGGCGCAAAA		
	CCCCAGTCCC	ACTACCTCCG	TCCGCGTTTT	TTCCTCCCAC	CAAACCOORDO
7851			CAACCCGAAA		
, 001	GTAGCACCAC	TTCCTCC A A C	GTTGGGCTTT	TTTGAGAACA	TTGCAGAAGG
7901					
7901	1"1"I'AAGAGC'I'	CTCCTGGCTA	GGAGTCACGT	AGAAAGGACT	ACCGACGAAG
			CCTCAGTGCA		
7951	GAACTTGGGT	CGCCGGTGTG	${\tt TTCGTATATG}$	GAGGTAGTAA	GACCTCCCTT
	CTTGAACCCA	GCGGCCACAC	AAGCATATAC	CTCCATCATT	CTGGAGGGAA
8001	TACAACCTAA	GGCGAGGAAC	TGCCCTTGCT	ATTCCACAAT	GTCGTCTTAC
	ATGTTGGATT	CCGCTCCTTG	ACGGGAACGA	TAAGGTGTTA	CAGCAGAATG
8051			TTGGAATGGC		
	TGGTAACTCA	GCAGAGGGGA	AACCTTACCG	GGGACCTGGG	CCGGGTGTTG
8101			ATTGTCTGTT		
	GACCGGGCGA	TTCCCTCAGG	TAACAGACAA	TAAAGTACCA	GAAAAATGTT
8151			TTTGAAGGAT		
	TGAGTATATA	AACGACTCCA	AAACTTCCTA	CCCTAATTCC	TCC ACA ATA
8201			GCAATATCAG		
	CTGTTTCGGG	CGAGGATGGA	CGTTATAGTC	CCACTCACAC	TGCAGCTTTG
8251			CCCTGGTTTC		
0251	TCCTACCTCA	TCTA A ACCCA	GGGACCAAAG	CACCTATGGT	GGAAGGGGCT
8301					
0301	CCCCCCCTCC	CACTACTICACGG	AGATGACGGA	GATGAAGGAG	GTGATGGAGA
0251			TCTACTGCCT		
8351	1GAGGGTGAG	GAAGGGCAGG	AGTGATGTAA	CTTGTTAGGA	GACGCCCTCA
			TCACTACATT		
8401	ATCGTATTAA	AAGCCGTGTA	TTCCCCCGCA	CTAAAGAATA	AATCCCCAGT
			AAGGGGGCGT		
8451	AGACATCATG	CGTGCTGTTG	GTGTATTTCT	GGCCATCTGT	CTTGTCACCA
	TCTGTAGTAC	GCACGACAAC	CACATAAAGA	CCGGTAGACA	GAACAGTGGT
8501	${\tt TTTTCGTCCT}$	CCCAACATGG	GGCAATTGGG	CATACCCATG	TTGTCACGTC
	AAAAGCAGGA	GGGTTGTACC	CCGTTAACCC	GTATGGGTAC	AACAGTGCAG
8551			CCTTCTCGCG		
	TGAGTCGAGG	CGCGAGTTGT	GGAAGAGCGC	AACCTTTTGT	AATCGCTGTA
8601			CATGCGACGG		
	AATGGACCAC	TCGTTAGTCT	GTACGCTGCC	GAAATCGGAC	CCCICCITAA
					COGROGRATT

8651			AACCAGCATG TTGGTCGTAC		
8701			AGGTGGCGGC		
0.01			TCCACCGCCG		
8751	CCACTCTACT	ACTGGGTATC	ATATGCTGAC	TGTATATGCA	TGAGGATAGC
	GGTGAGATGA	TGACCCATAG	TATACGACTG	ACATATACGT	ACTCCTATCG
8801	ATATGCTACC	CGGATACAGA	TTAGGATAGC	ATATACTACC	CAGATATAGA
	TATACGATGG	GCCTATGTCT	AATCCTATCG	TATATGATGG	GTCTATATCT
8851	${\tt TTAGGATAGC}$	ATATGCTACC	CAGATATAGA	TTAGGATAGC	CTATGCTACC
	AATCCTATCG	TATACGATGG	${\tt GTCTATATCT}$	AATCCTATCG	GATACGATGG
8901	CAGATATAAA	TTAGGATAGC	ATATACTACC	CAGATATAGA	TTAGGATAGC
	GTCTATATTT	AATCCTATCG	TATATGATGG	${\tt GTCTATATCT}$	AATCCTATCG
8951	ATATGCTACC	CAGATATAGA	TTAGGATAGC	CTATGCTACC	CAGATATAGA
	TATACGATGG	GTCTATATCT	AATCCTATCG	${\tt GATACGATGG}$	GTCTATATCT
9001			CAGATATAGA		
	AATCCTATCG	TATACGATGG	GTCTATATCT	AATCCTATCG	TATACGATAG
9051			GCTACCCAGA		
	GTCTATAAAC	CCATCATATA	CGATGGGTCT	ATATTTAATC	CTATCGTATA
9101			GATAGCATAT		
			CTATCGTATA		
9151			TATAGATTAG		
			ATATCTAATC		
9201			GCTACCCAGA		
			CGATGGGTCT		
9251			GATAGCATAT		
0201			CTATCGTATA		
9301			TATAGATTAG		
9351			ATATCTAATC		
3331			CCCATGGCAA GGGTACCGTT		
9401			GACCAACAAC		
7401			CTGGTTGTTG		
9451			CTCCAGATCG		
7 10 1			GAGGTCTAGC		
9501			GCAGGTATTC		
			CGTCCATAAG		
9551			CCGCAGTGGT		
			GGCGTCACCA		
9601			TACAGTCCAA		
			ATGTCAGGTT		
9651	GGGCTGACGC	GTGCCCCCAC	TCCACAATTT	CAAAAAAAAG	AGTGGCCACT
			AGGTGTTAAA		
9701	TGTCTTTGTT	TATGGGCCCC	ATTGGCGTGG	AGCCCCGTTT	AATTTTCGGG
	ACAGAAACAA	ATACCCGGGG	TAACCGCACC	TCGGGGCAAA	TTAAAAGCCC
9751			GAGTCCGCTG		
	CCACAATCTC	TGTTGGTCAC	CTCAGGCGAC	GACAGCCGCA	GGTGAGAGAA

### pCEP4W/hep36

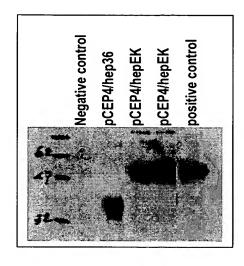
9801		ACAAATAGAG TGTTTATCTC			
9851	CTGCCTGGGA	CACATCTTAA	TAACCCCAGT	ATCATATTGC	ACTAGGATTA
9901		GTGTAGAATT ATAGCCATAA			
9951		TATCGGTATT CACCCCATGG			
10001	CCGAACAGGG	GTGGGGTACC ACTAGTATTT	TAAAGATAAC	AATTTCTATA	AGTCTTACAA
10001	AGTAAGGATG	TGATCATAAA	${\tt TAACGGGTTC}$	CCCAAACACT	CCCAATATAA
10051		CACAATGCCA GTGTTACGGT			
10101		CCTGAAACCT GGACTTTGGA			
10151		CAGCAGTTAT GTCGTCAATA			
10201	GCAGGCGAAG	ATTCAGGAGA	GTTCACTGCC	CGCTCCTTGA	TCTTCAGCCA
10251	CTGCCCTTGT	TAAGTCCTCT GACTAAAATG	GTTCACTACC	CTCGTGGAAT	CCTGACCCCA
10301		CTGATTTTAC ACCGTGACAG			
10351		TGGCACTGTC TACTAACCCT			
10401	TCCTGGGAAA	ATGATTGGGA TTGAATTAGG	${\tt TTAAGCTATC}$	GTATACGAAG	GGCAACCCAT
	TGTATACGAT	AACTTAATCC	CAATCAGACC	TATCATATAT	GATGATGGGC
10451	CCTTCGTATA	GCTACCCGTT CGATGGGCAA	ATCCCAATTG	TTCCCCCGGA	ATATTTGTGA
10501		CCCTCTTGAG GGGAGAACTC			
10551		GTTGGTGTAG CAACCACATC			
10601		CCCAGCATTG GGGTCGTAAC			
10651	AAAGGCAATG	TTGTGTTGCA AACACAACGT	GTCCACAGAC	TGCAAAGTCT	GCTCCAGGAT
10701	GAAAGCCACT	CAGTGTTGGC	AAATGTGCAC	ATCCATTTAT	AAGGATGTCA
10751	ACTACAGTCA	GTCACAACCG GAGAACCCCT	TTGTGTTTGG	TCCCCCCCG	TGTCACATGT
10801	GGAACAGGGC	CTCTTGGGGA CCAGTTGGCA	AGTTGTACCA	ACCAACTGAA	GGGATTACAT
10051		GGTCAACCGT			
10851		GCGAAGAAGG CGCTTCTTCC			
10901		GGGCTCTAGA CCCGAGATCT			

### FIGURE 11-10

### pCEP4W/hep36

10951	00000330300	0000000000			
10951			CCCTGACGGG		
			GGGACTGCCC		
11001	GCTTACAGAC	AAGCTGTGAC	CGTCTCCGGG	AGCTGCATGT	GTCAGAGGTT
			GCAGAGGCCC		
11051	TTCACCGTCA	TCACCGAAAC	GCGCGAGGCA	GCCGGATCAT	AATCAGCCAT
•	AAGTGGCAGT	AGTGGCTTTG	CGCGCTCCGT	CGGCCTAGTA	TTAGTCGGTA
11101			ACTTGCTTTA		
	TGGTGTAAAC	ATCTCCAAAA	TGAACGAAAT	TTTTTGGAGG	GGTGGAGGGG
11151	CTGAACCTGA	AACATAAAAT	GAATGCAATT	GTTGTTGTTA	ACTTGTTTAT
	GACTTGGACT	TTGTATTTTA	CTTACGTTAA	CAACAACAAT	TGAACAAATA
11201	TGCAGCTTAT	AATGGTTACA	AATAAAGCAA	TAGCATCACA	AATTTCACAA
	ACGTCGAATA	TTACCAATGT	TTATTTCGTT	ATCGTAGTGT	TTAAAGTGTT
11251			CATTCTAGTT		
	TATTTCGTAA	AAAAAGTGAC	GTAAGATCAA	CACCAAACAG	GTTTGAGTAG
11301			GATCCCACGT		
			CTAGGGTGCA		
11351			GAGGGCGAAG		
	TTCCCTCTAG	GCTGAGCAGA	CTCCCGCTTC	CGCTTCTGCG	CCTTCTCCGG
11401			CGGGAAGGAA		
	CGTCTCGGCC	GTCGTCCGGC	GCCCTTCCTT	CCAGGCGACC	TAACTCCCGG
11451			CGTCCCGCGC		
	CTTCCCTGCA	TCGTCTTCCT	GCAGGGCGCG	TCTTAGGTCC	ACCGTTGTGT
11501			ACGATGATTT		
	CCGCTCGTCG	GTTCCTTTCC	TGCTACTAAA	GGGGCTGTTG	TGGTGCCTTA
11551			GCCCCTGTCC		
	ACAGTCACGG	GTTGTCGGCT	CGGGGACAGG	TCGTCGCCCG	TTCCGTCCGC
11601			AATAGGGAGG		
	CGCTACTCAA	GGCGGCACCG	TTATCCCTCC	CCCTTTCGCT	TTCAGGGCCT
11651			CAATGCCCCA		
	TTCCTCGACT	GTCCACCACC	GTTACGGGGT	TGGTCACCCC	CAACGCAGTC
11701			CCACGTTGCC		
	GTTTGTGTCA	CGTGTGGTGC	GGTGCAACGG	ACTGTTGCCC	GGTGTTGAGG
11751			GGATTTATAC		
			CCTAAATATG		
11801			GATACAAAGG		
	GTATGCCCTT	CGTTATCGTA	CTATGTTTCC	GTAATTTCGT	CCCATACCTC
11851			TAGTTAAGAA		
	TATCGCATTT	TCCTCGTTGT	ATCAATTCTT	ATCCAGICAA	ACA A ACTORD
11901		CCAGAGGTTG		OGICAGII	MANAGEGE
•		GGTCTCCAAC	<del>-</del>		

Figure 12: Western Blot of Hepsin expressed in 293EBNA cells. 4 days after transfection, conditioned medium was harvested. 10  $\mu l$  of conditioned medium was loaded on a 10% Tris-Glycine gel. Untreated 293EBNA cells served as negative control, 5 ng of purified hepsin ED/EK (expressed from insect cells) served as positive control. After electrophoresis, proteins were transferred onto Nitrocellulose membrane at 100 V for 1h. The membrane was then probed with anti-V5 antibody at 37C° for 1h after blocking with 5% dry milk in TTBS buffer for 1h.



ਜੂ gnedium ₩W Anti-V5 antibody Eortrol cells

plague-purified clone 1

cells + medium plaque-purified clone 2 cells + medium

MW

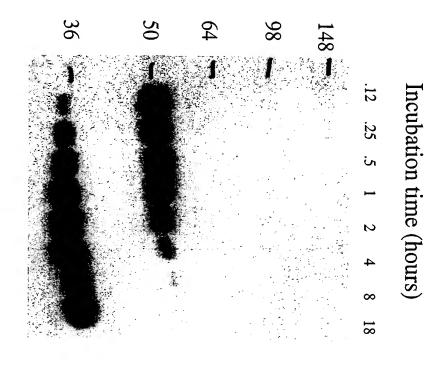
MW

Hopsin

Anti-66 48 33 25 1000 Anti-His antibody Countrol cells ្សាន្ត្បីue-purified clone 1 cells + medium plaque-purified clone 2 cells + medium

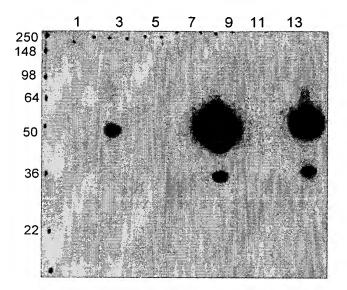
soluble hepsin was present in cultured medium and cell pellets. antibody (left panel) and an anti-His antibody (right panel) showed that recombinant was expressed in baculovirus infected insect cells. Western analysis using an anti-V5 Fig. 13. Western analysis of recombinant soluble hepsin. Recombinant soluble hepsin

Figure 14: Enterokinase (EKMax)processing of Hepsin -ED-EK protein to generate active enzyme



3.1 units EKMax / ml

Figure 15A: Western Slot Blot on the hybridomas 11C1 & 47A5 are WB positive against Hepsin-ED-EK protein.



Blot 1			
slot#	I.D. CODE		
1	1	2F10	
2	2	9H6	
3	3	11C1	
4	4	16A2	
5	5 27A1		
6	6 30C10		
7	7 30H12		
8	8 35D4		
9	9 47A5		
10	10 80F6		
11	Blank		
12	Blank		
13	Blank		
14	Mouse Poly clonal		

Figure 15B: Western Slot Blot on the hybridomas 31C1 and 38E2 are WB positive against Hepsin-ED-EK protein.

Blot 1			
slot#	I.D.	CODE	
1	1	7H3	
2	2	12A2	
3	3	16A3	
4	4	20D1	
5	5	22A3	
6	6	31C1	
7	7	31H2	
8	8	36B2	
9	9	38 E2	
10	10	39D6	
11	11	41F7	
12	12	42 E3	
13	Blank		
14	Mouse Poly clonal		

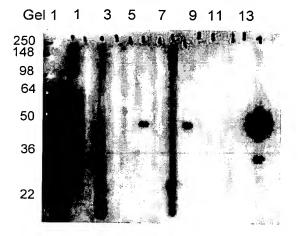
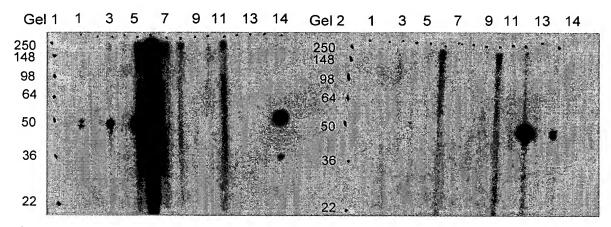


Figure 15C: Western Slot Blot on the hybridomas 37G10, 46D12 & 14C7 are WB positive against Hepsin-ED-EK protein.



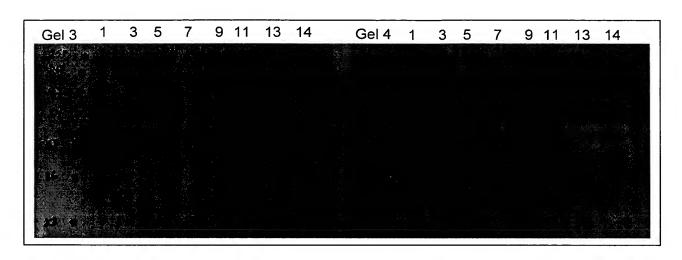
Gel 1

Slot	Sample	Dilution
1	Rabbit Polyclonal –Cayman	1:500
	Chem	
2	Media	neat
3	37G10	"
4	94A7	"
5	46D12	"
6	103E3	"
7	40F1	"
8	103H12	"
9	93D1	44
10	10C2	44
11	102F2	££
12	83E11	и
13	Blank	
14	Blank	
15	Protein - O mouse	1:5000

Gel 2

Slot	Sample	Dilution
1	Protein - O mouse	1:5000
2	Media	neat
3	27E7	и
4	92A7	44
5	91A4	"
6	99B11	44
7	94C7	и
8	91A1	"
9	14H11	11
10	74C7	"
11	72H6	и
12	14C7	и
13	Blank	
14	Rabbit Polyclonal –Cayman Chem	1:500

Figure 15D: Western Slot Blot on the hybridomas 14C7 & 72H6 are WB positive against Hepsin-ED-EK protein.



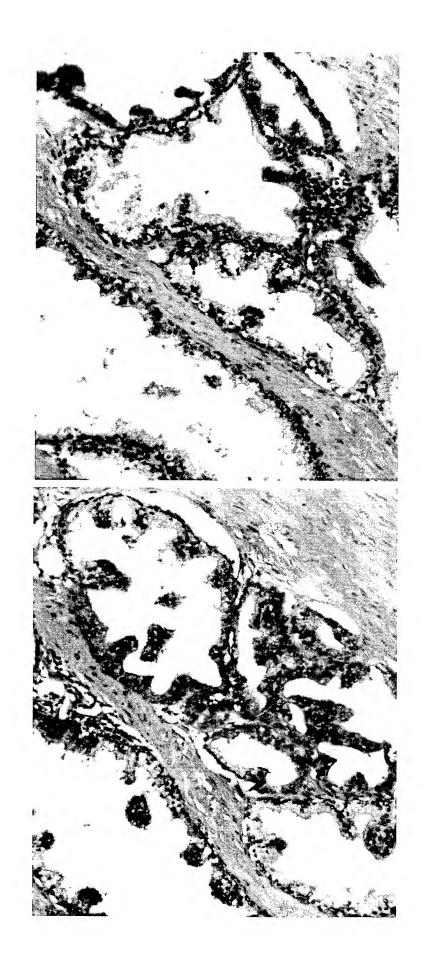
Gel 3

Slot	Sample	Dilution
1	Protein - 🔿 mouse	1:5000
2	Media	neat
3	84G6	66
4	84H2	66
5	51F8	tt
6	15b11	tt
7	90E6	"
8	72H6	и
9	85A4	"
10	87C2	11
11	3G11	11
12	53E11	66
13	91H4	66
14	53C7	a a
15	88C7	11

Gel 4

Slot	Sample	Dilution
1	Protein - O mouse	1:5000
2	Media	neat
3	75H3	"
4	98B4	"
5	91C9	"
6	53D9	"
7	80G6	"
8	95F3	"
9	14C7	íí.
10	80H10	íí.
11	92A9	íí.
12	81C8	u
13	96B6	a
14	1812	tt

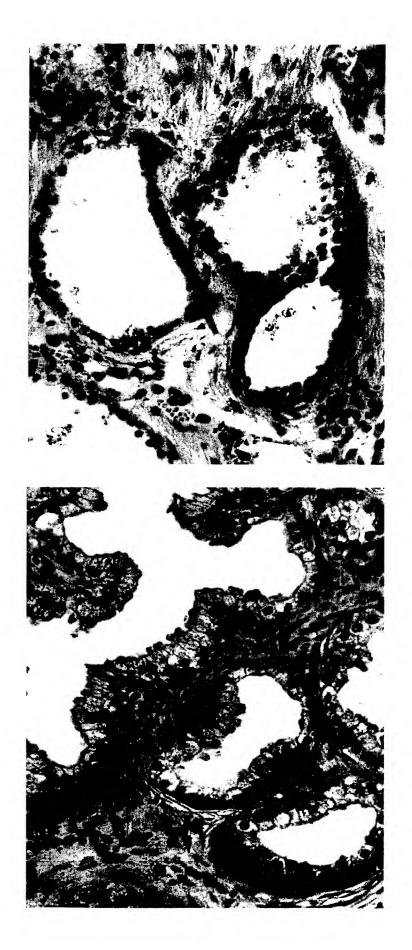
# FIG 16A: Human Prostate Tumor Tissue



Control
Anti-Hepsin mouse
polyclonal preimmun
1/500 (9)

Anti-Hepsin mouse polyclonal immun 1/500 (10)

# FIG 16B: Human Prostate Tumor Tissue



Control medium

Anti-Hepsin monoclonal ab (medium)

## FIG 16C: Human Prostate Tumor Tissue



Anti-Hepsin monoclonal ab

Figure 17

### Human wild-type hepsin amino acid sequence:

maqkeggrtv pccsrpkvaa ltagtllllt aigaaswaiv avllrsdqep lypvqvssad 61 arlmvfdkte gtwrllcssr snarvaglsc eemgflralt hseldvrtag angtsgffcv 121 degrlphtqr llevisvcdc prgrflaaic qdcgrrklpv drivggrdts lgrwpwqvsl 181 rydgahlcgg sllsgdwvlt aahcfpernr vlsrwrvfag avaqasphgl qlgvqavvyh 241 ggylpfrdpn seensndial vhlssplplt eyiqpvclpa agqalvdgki ctvtgwgntq 301 yygqqagvlq earvpiisnd vcngadfygn qikpkmfcag ypeggidacq gdsggpfvce 361 dsisrtprwr lcgivswgtg calaqkpgvy tkvsdfrewi fqaikthsea sgmvtql

the cytoplasmic domain: Met1 to Lys17 the transmembrane domain: Val18 to Leu44

the ectodomain: Arg45 to Leu417

Figure 18

Hep-ED-EK structure (modified soluble hepsin with substitute activation sequence)

```
1 RSDQEPLYPV QVSSADARLM VFDKTEGTWR LLCSSRSNAR VAGLSCEEMG FLRALTHSEL
61 DVRTAGANGT SGFFCVDEGR LPHTQRLLEV ISVCDCPRGR FLAAICQDCG RRKLPVDDDD
121 KIVGGRDTSL GRWPWQVSLR YDGAHLCGGS LLSGDWVLTA AHCFPERNRV LSRWRVFAGA
181 VAQASPHGLQ LGVQAVVYHG GYLPFRDPNS EENSNDIALV HLSSPLPLTE YIQPVCLPAA
241 GQALVDGKIC TVTGWGNTQY YGQQAGVLQE ARVPIISNDV CNGADFYGNQ IKPKMFCAGY
301 PEGGIDACQG DSGGPFVCED SISRTPRWRL CGIVSWGTGC ALAQKPGVYT KVSDFREWIF
361 QAIKTHSEAS GMVTQLEFGK PIPNPLLGLD STRTGHHHHH H*
```

Cytoplasmic domain: absent Transmembrane domain: absent

Modified activation domain:117-121 (underlined)

V5 and 6-His Tag: 377-401

Figure 19

### **Antibody Neutralization**

